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(54) Title: **METHOD**

(57) Abstract: The invention relates to a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or 2 as defined herein whereby to predict an increased likelihood of response to the erbB receptor drug. Preferred genes include any one of NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2, VAMP4, DAPK1, DAPK2, MLLT3, TNNC1, KIAA0931, ACOX2, EMP1, SLC20A1, SPRY2 or PGM1.

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## METHOD

The present invention relates to sensitivity of tumours to therapeutic agents which can be predicted from the gene expression profile of the tumour and hence that the suitability of cancer patients for treatment with such therapeutic agents can be determined by measuring the relative expression levels of particular genes in tumour tissue.

The phosphorylation of proteins on tyrosine residues is a key element of signal transduction within cells. Enzymes capable of catalysing such reactions are termed tyrosine kinases. A number of these enzymes exist as integral components of transmembrane receptor molecules and are classified as receptor tyrosine kinases (RTKs). There are several members of this family of RTKs, class I of which includes the erbB family, e.g. epidermal growth factor receptor (EGFR), erbB2, erbB3 and erbB4. Binding of a variety of ligands to the external domain activates the EGFR tyrosine kinase domain. Activation causes EGFR itself and a number of cellular substrates to become phosphorylated on tyrosine residues. These phosphorylation reactions are a major component of growth factor induced proliferation of cells.

The erbB family of receptor tyrosine kinases are known to be frequently involved in driving the proliferation and survival of tumour cells (reviewed in Olayioye *et al.*, *EMBO J.*, 2000, 19, 3159). One mechanism by which this can occur is over expression of the receptor at the protein level, for example as a result of gene amplification. This has been observed in many common human cancers (reviewed in Klapper *et al.*, *Adv. Cancer Res.*, 2000, 77, 25) such as, non-small cell lung cancers (NSCLCs) including adenocarcinomas (Cerny *et al.*, *Brit. J. Cancer*, 1986, 54, 265; Reubi *et al.*, *Int. J. Cancer*, 1990, 45, 269; Rusch *et al.*, *Cancer Research*, 1993, 53, 2379; Brabender *et al.*, *Clin. Cancer Res.*, 2001, 7, 1850) as well as other cancers of the lung (Hendler *et al.*, *Cancer Cells*, 1989, 7, 347).

It is now several decades since the study of retroviral mediated cellular transformation began to revolutionize our understanding of malignant transformation. Transformation was shown to be dependent on oncogenes carried by viruses and these were shown to have mammalian cellular counterparts, proto-oncogenes. In 1984, EGFR was described as the mammalian counterpart of the retroviral oncogene, v-erbB (Downward *et al.*). This, coupled to earlier observations describing a two component autocrine growth promoting mechanism in cancer cells consisting of EGF ligand and its receptor EGFR (Sporn & Todaro), strengthened

the hypothesis that EGFR signalling is an important contributor to tumourigenesis. Subsequent reports continued to provide evidence that EGFR is an attractive target for therapeutic intervention in Cancer (see Yarden & Sliwkowski for review). EGFR is markedly overexpressed across a large variety of epithelial Cancers (see Salomon et al) and some immunohistochemical studies have demonstrated EGFR expression is associated with poor prognosis. In addition to overexpression, it is recognised that there is potential for deregulated EGFR signalling in tumours via a number of alternative mechanisms including i) EGFR mutations ii) increased ligand expression and enhanced autocrine loop and iii) heterodimerisation and cross talk with other erbB receptor family members.

In addition, a wealth of pre-clinical information suggests that the erbB family of receptor tyrosine kinases are involved in cellular transformation. In addition to this, a number of pre-clinical studies have demonstrated that anti-proliferative effects can be induced by knocking out one or more erbB activities by small molecule inhibitors, dominant negatives or inhibitory antibodies (reviewed in Mendelsohn et al., Oncogene, 2000, 19, 6550).

Thus it has been recognised that inhibitors of these receptor tyrosine kinases should be of value as a selective inhibitor of mammalian cancer cells (Yaish et al. Science, 1988, 242, 933, Kolibaba et al, Biochimica et Biophysica Acta, 1997, 133, F217-F248; Al-Obeidi et al, 2000, Oncogene, 19, 5690-5701; Mendelsohn et al, 2000, Oncogene, 19, 6550-6565).

A number of small molecule inhibitors of erbB family of receptor tyrosine kinases are known, particularly inhibitors of EGF and erbB2 receptor tyrosine kinases. For example European Patent Application No. 0566226 and International Patent Applications WO 96/33980 and WO 97/30034 disclose that certain quinazoline derivatives which possess an anilino substituent at the 4-position possess EGFR tyrosine kinase inhibitory activity and are inhibitors of cancer tissue.

It has been disclosed by J R Woodburn et al. in Proc. Amer. Assoc. Cancer Research, 1997, 38, 633 and Pharmacol. Ther., 1999, 82, 241-250 that the compound N-(3-chloro-4-fluorophenyl)-7-methoxy-6-(3-morpholinopropoxy)quinazolin-4-amine is a potent EGFR tyrosine kinase inhibitor. This compound is also known as Iressa (registered trade mark), gefitinib (United States Adopted Name), by way of the code number ZD1839 and Chemical Abstracts Registry Number 184475-35-2. The compound is principally identified hereinafter as gefitinib.

Gefitinib was developed as an inhibitor of epidermal growth factor receptor-tyrosine kinase (EGFR-TK), which blocks signalling pathways responsible for driving proliferation, invasion, and survival of cancer cells (Wakeling, A.E., et al. *Cancer Res*, 2002, 62(20), p5749). Gefitinib has provided clinical validation of small molecule inhibitors of EGFR. Potent anti-tumour effects as well as rapid improvements in NSCLC-related symptoms and quality of life have been observed in clinical studies that enrolled patients with advanced NSCLC who did not respond to platinum-based chemotherapy. The Phase II 'IDEAL' trials demonstrated that single agent gefitinib resulted in objective anti-tumour activity, symptomatic improvement and limited toxicity in patients with advanced NSCLC and previously treated with cytotoxic chemotherapy (Fukuoka et al., Kris et al). Objective response rate (Complete Response + Partial Response) was 18.4% and 11.8% respectively in the IDEAL 1 and IDEAL 2 trials. The differences in response in these clinical trials has been attributed to different population groups in the two trials, predominantly Japanese in IDEAL 1 and a predominantly European-derived population in IDEAL 2. Beyond objective responses, additional patients experienced stable disease and / or symptom improvement meaning that approximately 50% of patients overall benefit from gefitinib. The tumour response data has been the basis of initial regulatory approvals of gefitinib in advanced NSCLC in several markets.

It is important to be able to understand the basis of response to anti-cancer therapeutic agents such as gefitinib since this would allow clinicians to maximise the benefit/risk ratio for each patient, potentially via the development of diagnostic tests to identify patients most likely to benefit from gefitinib treatment. An obvious candidate marker of response to gefitinib has been EGFR expression level. However, gefitinib inhibition of growth of some cancer-derived cell lines and tumour xenografts is not well correlated with the level of expression of EGFR. Furthermore, studies alongside the IDEAL trials demonstrated that EGFR protein expression as measured by IHC was not an accurate predictor of response to gefitinib (Bailey et al). Although there are now several additional hypotheses based on genetics, genomics, proteomics, biochemical and other studies, there is still no pre-treatment predictive biomarker of gefitinib response currently approved by regulatory authorities. Possibly the most significant recent breakthrough in understanding gefitinib response has come from recent data (Lynch et al, Paez et al) indicating that mutation in the EGFR kinase domain predicts gefitinib hypersensitivity in NSCLC patients. Hypersensitivity is a vague term but in this field is generally understood to mean patients experiencing objective tumour responses (i.e. marked tumour regression,

normally above 50%). As well as demonstrating the EGFR mechanism of action for gefitinib, this may provide a basis for venturing into other disease settings such as first line, adjuvant and possibly earlier cancer intervention with EGFR inhibitors in a targeted subpopulation in NSCLC patients and other types of cancers carrying the EGFR mutation.

However, it is likely that restricting prescription of gefitinib to the mutant EGFR carrying tumour subgroup will deprive many patients who could benefit from gefitinib. Firstly there are emerging reports of gefitinib hypersensitive patients with undetectable EGFR mutation in their tumour and other patients with EGFR mutation who do not respond to gefitinib. Secondly, data reported at ASCO 2004 (Shepherd et al) indicated that the EGFR small molecule tyrosine kinase inhibitor erlotinib (Roche, Genentech, OSI) prolongs survival in advanced NSCLC previously treated with chemotherapy, by ~2 months across the population with resulting 41% reduction in risk of death at one year. Most interestingly, the survival benefit appears to be derived from patients in the stable disease response population as well as hypersensitive patients. This highlights the likely importance of identifying likely gefitinib responsive patients beyond those carrying EGFR mutation. Definitive survival benefit is also likely to be demonstrated from ongoing clinical trials with gefitinib.

The differential response of patients to chemotherapy treatments indicates that there is a need to find methods of predicting which treatment regimes best suit a particular patient.

There is an increasing body of evidence that suggests that patients' responses to numerous drugs may be related to a patients' genetic, genomic, proteomic, biochemical or profile and that determination of the genetic factors that influence, for example, response to a particular drug could be used to provide a patient with a personalised treatment regime. Such personalised treatment regimes offer the potential to maximise therapeutic benefit to the patient, whilst minimising, for example side effects that may be associated with alternative and less effective treatment regimes.

Therefore there is a need for methods that can predict a patients' response to a drug based on the results of a test that indicates whether the patient is likely to respond to treatment or to be resistant to treatment.

The present invention is based on the discovery that the sensitivity of tumours to therapeutic agents can be predicted from the gene expression profile of the tumour and hence that the suitability of tumour patients for treatment with such therapeutic agents can be determined by measuring the relative expression levels of particular genes in tumour tissue.

According to one aspect of the present invention there is provided a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 as defined herein whereby to predict an increased likelihood of response to the erbB receptor drug.

According to another aspect of the present invention there is provided a method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or DAPK2 whereby to predict an increased likelihood of response to the erbB receptor drug.

In one embodiment the method comprises testing a biological sample from the mammal for expression of any one of ACOX2, NPAS2, NES, CHST7, GSPT2, DAPK1, DAPK2 or TNNC1. More preferably the method comprises testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7 or DAPK1. More preferably the method comprises testing a biological sample from the mammal for expression of at least two of NPAS2, NES, CHST7 or DAPK1. More preferably the method comprises testing a biological sample from the mammal for expression of at least three of NPAS2, NES, CHST7 or DAPK1. More preferably still the method comprises testing a biological sample from the mammal for expression of NPAS2, NES, CHST7 and DAPK1.

In an alternative embodiment the method comprises testing a biological sample from the mammal for expression of any one of NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2, VAMP4, DAPK1, DAPK2, MLLT3, TNNC1 or KIAA0931. More preferably the method comprises testing a biological sample from the mammal for expression of any one of DAPK1, DAPK2 or NES. More preferably the method comprises testing a biological sample from the mammal for expression of at least two of DAPK1, DAPK2 or NES. More preferably the method comprises testing a biological sample from the mammal for expression of DAPK1, DAPK2 and NES.

In a preferred embodiment the method additionally comprises testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1, SLC20A1, SPRY2 or PGM1. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an alternative preferred embodiment the method additionally comprises testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1, HCA127, UBL5, ZNF23, UROD, CD44, SPRY1, RAPGEF2, SLC20A1, NRP1, PGM1, SPRY2, PTGER3, SCN10A, KITLG, CDH1, HOP, BCL3 or OLFM1. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

Preferably the tumour is selected from the group consisting of leukaemia, multiple myeloma, lymphoma, bile duct, bone, bladder, brain, CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head, neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural membrane, peritoneal membrane, prostate, renal, skin, testicular, thyroid, uterine and vulval. More preferably the tumour is selected from one of non-small cell lung, pancreatic, head or neck. More preferably the tumour is selected from one of non-small cell lung, head or neck.

Preferably the erbB receptor drug is selected from any one of gefitinib, erlotinib, PKI-166, EKB-569, HKI-272, lapatinib, canertinib, AEE788, XL647, BMS 5599626, cetuximab, matuzumab, panitumumab, MR1-1, IMC-11F8 or EGFR11. Most preferably the erbB receptor drug is gefitinib.

In a further preferred embodiment of the method of the invention the mammal is a human and the method comprises testing a biological sample from the human for increased expression of DAPK1 and decreased expression of NPAS2, NES, CHST7 or EMP1 whereby to predict an increased likelihood of response to gefitinib. In an alternative preferred embodiment of the method of the invention the mammal is a human and the method comprises testing a biological sample from the human for increased expression of DAPK1 and DAPK2 and decreased expression of NES and EMP1 whereby to predict an increased likelihood of response to gefitinib.

According to another aspect of the invention there is provided an isolated set of marker genes identified as having differential expression between tumour cells that are sensitive and resistant to an erbB receptor drug said gene set comprising one or more genes selected from at least the group consisting of the genes listed in Table 1 defined herein or DAPK2, including gene specific oligonucleotides derived from said genes. Preferably the set comprises at least 2



genes, more preferably at least 3 genes, more preferably at least 4 genes. More preferably the set comprises at least one gene selected from Table 2 as defined herein.

According to another aspect of the invention there is provided an isolated set of marker genes identified as having differential expression between tumour cells that are sensitive and resistant to an erbB receptor drug said gene set comprising one or more genes selected from at least the group consisting of the genes listed in Table 1 defined herein, including gene specific oligonucleotides derived from said genes. Preferably the set comprises at least 2 genes, more preferably at least 3 genes. More preferably the set comprises at least one gene selected from Table 2 as defined herein.

The present invention permits the improved selection of a patient, having or suspected of having a tumour, for treatment with an erbB receptor drug, in order to predict an increased likelihood of response to the erbB receptor drug.

In one embodiment, the method comprises testing a biological sample from the mammal for expression of at least one or more of the following from Table 1, which are found at lower levels in sensitive cells NPAS2, NES, CHST7, ACOX2 or GSPT2 or at least one or more of the following which are found at higher levels in sensitive cells DAPK1 or TNNC1. The Affymetrix ID and Affymetrix probe sequence for these genes are displayed in Table 1. In a preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of DAPK2 which is found at higher levels in sensitive cells, whereby to predict an increased likelihood of response to the erbB receptor drug.

In an alternative embodiment, the method comprises testing a biological sample from the mammal for expression of at least one or more of the following from Table 1, which are found at lower levels in sensitive cells NES, GSPT2, ETR101, TAZ, CHST7, DNAJC3, NPAS2, PIN1, TCEA2 or VAMP4 or at least one or more of the following which are found at higher levels in sensitive cells DAPK1, DAPK2, MLLT3, TNNC1 or KIAA0931. The Affymetrix ID and Affymetrix probe sequence for these genes are displayed in Table 1.

In a preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 2, whereby to predict an increased likelihood of response to the erbB receptor drug. In a preferred embodiment, the method comprises testing a biological sample from the mammal for expression of any one of the following genes listed in Table 2, which are found at lower levels in sensitive cells EMP1, SLC20A1, SPRY2 or PGM1, whereby to predict an increased likelihood of response to the

erbB receptor drug. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an alternative preferred embodiment, the method further comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 2, whereby to predict an increased likelihood of response to the erbB receptor drug. In a preferred embodiment, the method comprises testing a biological sample from the mammal for expression of any one of the following genes listed in Table 2, which are found at lower levels in sensitive cells EMP1, HCA127, UBL5, ZNF23, UROD, CD44, SPRY1, RAPGEF2, SLC20A1, NRP1, PGM1 or SPRY2 or at least one or more of the following which are found at higher levels in sensitive cells PTGER3, SCN10A, KITLG, CDH1, HOP, BCL3 or OLFM1 whereby to predict an increased likelihood of response to the erbB receptor drug. More preferably the method comprises testing a biological sample from the mammal for expression of EMP1.

In an especially preferred embodiment the method comprises testing a biological sample from the mammal for expression of NPAS2, NES, CHST7, DAPK1 and EMP1. High NPAS2, NES, CHST7 and EMP1 levels are associated with resistance to gefitinib and high DAPK1 levels are associated with sensitivity to gefitinib. Preferably, the assessment of expression comprises determination of whether DAPK1 levels are increased and NPAS2, NES, CHST7 and EMP1 levels are reduced.

In an alternative especially preferred embodiment the method comprises testing a biological sample from the mammal for expression of DAPK1, DAPK2, NES and EMP1. High EMP1 and NES levels are associated with resistance to gefitinib and high DAPK1 and DAPK2 levels are associated with sensitivity to gefitinib. Preferably, the assessment of expression comprises determination of whether DAPK1 and DAPK2 levels are increased and EMP1 and NES levels are reduced. In a most preferred embodiment the invention comprises determining the level of DAPK1 and EMP1.

According to another aspect of the invention there is provided a method for predicting clinical outcome of treatment with an erbB receptor drug for a mammal, having or suspected of having a tumour, comprising determining the level of any of the genes as described hereinabove in a biological sample taken from the tumour, or suspected tumour, wherein a poor outcome is predicted if:

- a) the expression level of DAPK1 is reduced; and /or

- b) the expression level of NPAS2, NES, CHST7 and EMP1 is increased.

According to another aspect of the invention there is provided a method for classifying cancer comprising, determining the level of any of the genes as described hereinabove in a biological sample taken from a tumour, or suspected tumour, wherein tumours expressing elevated levels of DAPK1 and / or reduced levels of NPAS2, NES, CHST7 or EMP1 are predicted as sensitive to treatment with erbB receptor drugs.

According to another aspect of the invention there is provided a method for predicting clinical outcome of treatment with an erbB receptor drug for a mammal, having or suspected of having a tumour, comprising determining the level of any of the genes as described hereinabove in a biological sample taken from the tumour, or suspected tumour, wherein a poor outcome is predicted if:

- a) the expression level of DAPK1 or DAPK2 is reduced; and /or
- b) the expression level of EMP1 or NES is increased.

According to another aspect of the invention there is provided a method for classifying cancer comprising, determining the level of any of the genes as described hereinabove in a biological sample taken from a tumour, or suspected tumour, wherein tumours expressing elevated levels of DAPK1 or DAPK2 and / or reduced levels of EMP1 or NES are predicted as sensitive to treatment with erbB receptor drugs.

According to another aspect of the invention there is provided a method for treating a disease condition in a mammal having, or suspected of having, a tumour, predicted to be resistant or non responsive to erbB receptor drug treatment based on the level of any of the genes as described hereinabove, comprising: providing a resistance-surmounting quantity of an erbB receptor drug and administering the resistance-surmounting quantity of the erbB receptor drug to the mammal.

In a preferred embodiment the mammal is a primate. In a most preferred embodiment the mammal is a human. In a preferred embodiment the patient is a primate. In a most preferred embodiment the patient is a human.

The term "erbB receptor drug" includes drugs acting upon the erbB family of receptor tyrosine kinases, which include EGFR, erbB2 (HER), erbB3 and erbB4 as described in the background to the invention above. In a preferred embodiment the erbB receptor drug is an erbB receptor tyrosine kinase inhibitor. In a preferred embodiment the erbB receptor drug is an EGFR tyrosine kinase inhibitor.

In a more preferred embodiment the EGF receptor tyrosine kinase inhibitor is selected from gefitinib, Erlotinib (OSI-774, CP-358774), PKI-166, EKB-569, HKI-272 (WAY-177820), lapatinib (GW2016, GW-572016), canertinib (CI-1033, PD183805), AEE788, XL647, BMS 5599626 or any of the compounds as disclosed in WO03/082831, WO05/012290, WO05/026157, WO05/026150, WO05/026156, WO05/028470, WO05/028469, WO2004/006846, WO03082831, WO03/082290 or PCT/GB2005/000237.

In another preferred embodiment the erbB receptor drug is an anti-EGFR antibody such as for example one of cetuximab (C225), matuzumab (EMD-72000), panitumumab (ABX-EGF/rHuMAb-EGFr), MR1-1, IMC-11F8 or EGFR11.

We contemplate that erbB receptor drugs may be used as monotherapy or in combination with other drugs of the same or different classes. In an especially preferred embodiment the EGF receptor tyrosine kinase inhibitor is gefitinib.

In a preferred embodiment the present invention is particularly suitable for use in predicting the response to the erbB receptor drug as described hereinbefore in those patients or patient population with a tumour which is dependent alone, or in part, on an erbB tyrosine kinase receptor. Such tumours include, for example, non-solid tumours such as leukaemia, multiple myeloma or lymphoma, and also solid tumours, for example bile duct, bone, bladder, brain/CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head and neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural/peritoneal membranes, prostate, renal, skin, testicular, thyroid, uterine and vulval tumours.

In a preferred embodiment the present invention is particularly suitable for identifying a patient with head, neck, pancreatic, glioblastoma, colorectal or breast tumour for drug treatment. In an especially preferred embodiment the present invention also is particularly suitable for identifying those patients with NSCLC, more particularly advanced NSCLC including advanced adenocarcinoma that will respond to treatment with an erbB receptor drug as hereinbefore defined.

The present invention provides advantage in the treatment of tumours such as NSCLC, especially advanced NSCLC by identifying "individual cancer profiles" of NSCLC and so determining which tumours would respond to erbB receptor drug such as gefitinib.

The present invention is particularly useful in the treatment of patients with advanced NSCLC who have failed previous chemotherapy, such as platinum-based chemotherapy. The present invention is also particularly useful in the treatment of patients with locally advanced

(stage IIIB) or metastasized (stage IV) NSCLC who have received previous chemotherapy, such as platinum-based chemotherapy. The present invention is also useful in adjuvant therapy or as a first-line therapy.

In a preferred embodiment there is provided a method of selecting a human, having or suspected of having a tumour, for treatment with gefitinib which comprises testing a biological sample, from the mammal for expression of NPAS2, NES, CHST7, DAPK1 and EMP1, whereby to predict an increased likelihood of response to gefitinib.

In a preferred embodiment there is provided a method of selecting a human, having or suspected of having a tumour, for treatment with gefitinib which comprises testing a biological sample, from the mammal for expression of DAPK1, DAPK2, NES and EMP1 whereby to predict an increased likelihood of response to gefitinib.

According to another aspect of the invention there is provided a method of predicting the responsiveness of a patient or patient population with cancer, for example lung cancer, to treatment with chemotherapeutic agents, especially erbB receptor drugs, comprising comparing the differential expression of any of the genes described herein.

In one embodiment the assessment of expression is performed by gene expression profiling using oligonucleotide-based arrays or cDNA-based arrays of any type, particularly where large numbers of genes are analysed simultaneously. In an alternative embodiment, RT-PCR (reverse transcription- Polymerase Chain Reaction), real-time PCR, *in-situ* hybridisation, Northern blotting, Serial analysis of gene expression (SAGE) for example as described by Velculescu et al Science 270 (5235): 484-487, or differential display or any other method of measuring gene expression at the RNA level could be used. Details of these and other general molecular biology techniques can be found in Current Protocols in Molecular Biology Volumes 1-3, edited by F M Asubel, R Brent and R E Kingston; published by John Wiley, 1998 and Sambrook, J. and Russell, D.W., Molecular Cloning: A Laboratory Manual, the third edition, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York, 2001.

In another embodiment the assessment of expression is performed by measurement of protein levels encoded by the aforementioned genes. For example, an immunohistochemistry-based assay or application of an alternative proteomics methodology.

In another embodiment the assessment of expression is performed by measurement of activity of the proteins encoded by the aforementioned genes, for example in a bioassay.

In a preferred embodiment the biological sample would have been obtained using a

minimally invasive technique to obtain a small sample of tumour, or suspected tumour, from which to determine gene expression profile. Such techniques include, for example tumour biopsy, such as transbronchial biopsy. The profile of gene expression of transbronchial biopsy specimens whose size is about 1 mm may be measured for example using a suitable amplification procedure.

Another aspect of the invention provides a kit for use in a method of predicting the responsiveness of a patient or patient population with a tumour, to treatment with chemotherapeutic agents, especially erbB receptor drugs, comprising a means for measuring the levels of any of the genes as described hereinabove. Preferably the genes are attached to a support material or membrane such as nitrocellulose, or nylon or a plastic film or slide.

In a further preferred embodiment the present invention includes administration of an erbB receptor drug to a mammal selected according the methods described hereinabove.

According to another aspect of the invention there is provided a method of using the results of the methods described above in determining an appropriate dosage of an erbB receptor drug.

In a preferred embodiment the biological sample comprises either a single sample which may be tested for expression of any of the genes as described hereinabove, or multiple samples which may be tested for expression of one or more of the genes as described hereinabove.

The invention is illustrated by the following non-limiting examples in which:

Fig 1 illustrates a xenograft (A549 cell line) which when grown as a xenograft in athymic mice is sensitive to gefitinib. This involved oral dosing, once daily, at the dose indicated. Y axis = mean tumour volume in  $\text{cm}^3$ ; x axis = days after treatment.

Fig 2 illustrates a xenograft (MKN45 cell line) which when grown as a xenograft in athymic mice is resistant to gefitinib. This involved oral dosing, once daily, at the dose indicated. Y axis = mean tumour volume in  $\text{cm}^3$ ; x axis = days after treatment.

Figures 3, 4, 5 and 6 show examples of specific gene expression profiled across a wider panel of gefitinib sensitive and resistant lines, where definition of sensitivity is based on response to gefitinib when grown as a xenograft, to increase confidence that the expression profile of each gene is truly predictive. Iressa sensitivity is based on xenografts data. The cell lines and the tumours from which they are derived are as follows; KB – head and neck, HT29 - colon, BT474 – breast, DU145 – prostate, LoVo – colon, MCF7 – breast, GEO – colon, A549 – lung,

A431 - epidermoid, H322 - lung, HX147 - lung, RT112 - bladder, MiaPaCa2 - pancreas, MKN45 - gastric, MDAMB231 - breast, PC3 - prostate, Calu6 - lung, SW620 - colon.

The legend key is S=sensitive, U=unknown and R=resistant.

Fig 3 shows EMP1 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 4 shows DAPK1 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 5 shows DAPK2 basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

Fig 6 shows NES basal expression in Cell Culture - wider cell panel (Taqman RT-PCR).

### Example 1

#### **Gene Expression in Gefitinib Resistant or Sensitive Tumour Cell Lines – Cell Culture and Xenograft Studies**

We identified genes useful to predict response to erbB receptor drugs in the clinic. This is based on studies with gefitinib, but the findings are applicable to erbB receptor drugs in general.

The gene lists have been assembled by comparing tumour cell lines which have been demonstrated to be either sensitive to gefitinib or resistant to gefitinib. This definition is based on the response observed when the tumour cell line is implanted into nude mice and grown as a xenograft. This definition has been used for all the pre-clinical studies described herein.

Initially a small panel of six human tumour cell lines were assembled, three which are sensitive to gefitinib and three which are resistant to gefitinib in the xenograft setting defined above.

The sensitive cell lines were;

1. Lovo (ATCC<sup>1</sup> No. CCL-229) – colon tumour cell line
2. KB (ATCC No. CCL-17) – initially reported as a nasopharyngeal cell line (although more recently reported as Hela derived (cervical carcinoma))
3. HT29 (ATCC No. HTB-38) – colon tumour cell line

The resistant cell lines were;

1. MKN 45 (source - Nottingham University, UK) – gastric tumour cell line
2. Calu 6 (ATCC No. HTB-56) – lung tumour cell line
3. PC3 (ATCC No. CRL-1435) – prostate tumour cell line

<sup>1</sup>ATCC = American Type Culture Collection

The cell lines were grown both in cell culture and as xenografts, RNA prepared and the basal expression profiles determined by measuring RNA expression on the Affymetrix microarray platform. As part of our studies, the term 'basal' has been used to indicate constitutive or steady state expression levels (rather than expression levels which are modulated as a consequence of administration of an erbB ligand or gefitinib to the cells). Figure 1 illustrates the sensitivity of A549 xenografts (used in Example 3 below) to treatment with gefitinib. Figure 2 illustrates the resistance of MKN45 xenografts to gefitinib. See Example 2 below for analysis of results.



### Example 2

#### **Statistical analyses of cell culture and xenograft data sets**

The following statistical analyses were performed separately for cell culture and xenograft data sets. Probe sets were eliminated if their signal was not distinguishable from background noise across all RNA samples in the set. Mixed ANOVA (see for example Scheffe, 1959) was applied separately to each remaining probe set to generate p values. The p values were then used to calculate Q values (Storey). The Q values indicate the expected proportion of genes in a gene list which are not truly differentially expressed but have been falsely discovered (False Discovery Rate or FDR). Q value cut-offs appropriate in the different studies were identified and applied, based on graphical examination of the p value and Q value results, in conjunction with fold change. The final genelists for each study were generated using Q value and fold change (FC) cut-offs. The different genelists were then combined to display an overall list of genes which showed consistent differences in expression profiles between the cell lines in the sensitive and resistant groups.

Further details of the analysis procedures are provided as follows. Fold change (FC) was calculated based on the mean of sensitive cells divided by the mean of resistant cells. To generate gene lists, FC cut-off of two-fold (2X) change in either direction was used in all cases. Furthermore FDR Q values were used to narrow down the lists and obtain the most significant gene changes across sensitive versus resistant cell lines. In the case of cell culture, Q value cut-off is 0.3. In the case of xenograft, Q value cut-off is 0.6. The different cut-offs used reflect the different design and variance values associated with each experiment.

In cell culture studies, lists were obtained based on the above criteria for cells grown either in full serum containing medium or in charcoal stripped serum. In the xenograft study, the same as above was performed for separate sets of tumours harvested at 18hr intervals. Gene lists contain some redundancy in genes where appropriate to illustrate consistency of results obtained for example with different probe sets.

### Example 3

#### **Identification of predictive genes**

Genes which have not previously been identified as predictive of erbB receptor drug sensitivity are listed in Table 1. Other genes which we have identified to be optionally used in combination with Table 1 genes are listed in Table 2.

Key to Tables:

'Affymetrix ID' – the Affymetrix probe set identifier

'Sequence' – target sequence relating to the Affymetrix probe set indicated by 'Affymetrix ID'

"+" if up in sensitive" means that the gene is relatively highly expressed in sensitive cells. (Consequently, absence of a "+" means that the gene is relatively highly expressed in resistant cells).

'Gene Title' - The current annotation of the gene relating to 'Affymetrix ID' based on UniGene  
133

'Gene Symbol' – shorthand synonym for the gene title

'Locus Link' & RefSeq Transcript ID' are provided for gene identification purposes.

Combining genes has the potential to generate an improved diagnostic over genes used in isolation. Collective gene expression profiles (at the RNA and/ or protein level) may be more likely to identify patients most likely to benefit from gefitinib rather than the expression level of one gene in isolation.

It may be more practical when developing a pre-treatment response prediction diagnostic to work with a truncated gene list from tables 1 and / or 2. A number of criteria have been used to shorten the gene list to identify those genes which are most predictive of response. Firstly the statistical (p values and Q values or FDR values) can indicate the statistical significance of a gene.

Secondly, the differential expression (fold change) between the sensitive and resistant groups indicates the potential sensitivity of a marker to be used in a diagnostic test (highest fold change between sensitive group and resistant group is preferred).

Thirdly, we have performed RT-PCR based expression profiling across a wider panel of gefitinib sensitive and resistant human tumour cell lines to increase confidence that the expression profile of each gene is truly predictive. Figs 3, 4, 5 and 6 show examples of specific gene expression profiled across a wider panel of cell lines as set out below.

The sensitive human tumour cell lines, where definition of sensitivity is based on response to Iressa when grown as a xenograft:

- a. BT474 (ATCC No. HTB-20) – breast tumour cell line
- b. DU145 (ATCC No. HTB-81) – colon tumour cell line

- c. MCF7 (ATCC No. HTB-22, sourced from ICRF (now CR-UK), London), - breast tumour cell line
- d. GEO colon tumour cell line. RNA obtained from Fortunato Ciardiello, Cattedra di Oncologia Medica, Dipartimento Medico-Chirurgico di Internistica Clinica e Sperimentale "F. Magrassi e A. Lanzara, " Seconda Universita delgi Studi di Napoli, Via S. Pansini, 5-80131, Naples, Italy.
- e. A549 (ATCC No. CCL-185) – lung tumour cell line
- f. A431 (ATCC No. CRL-155) – epidermoid cell line

The resistant human tumour cell lines, where definition of sensitivity is based on response to Iressa when grown as a xenograft:

- 1) HX147 - (source: ICRF (now CR-UK), London) – lung tumour cell line
- 2) RT112 - bladder tumour cell line (DSMZ No ACC 418)
- 3) MiaPac2 (ECACC 85062806, ref. no. 001611) pancreatic tumour cell line
- 4) MDAMB231 (ATCC No. HTB-26) – breast tumour cell line
- 5) SW620 (ECACC CCL-227) – colon tumour cell line

ATCC = American Type Culture Collection

DSMZ - Deutsche Sammlung von Mikroorganismen und Zellkulturen GmbH (German Collection of Micro-organisms and Cell Cultures)

ECACC = European Collection of Cell Cultures

In isolation, each of these genes is reasonably predictive of gefitinib response, but collectively they can be applied to make predictions with a higher level of confidence.

The Affymetrix probe sets identifiers for the genes in the above diagnostic genelists are indicated in Tables 1 and 2. Current Affy IDs are based on Affy U133 chipset. For the avoidance of doubt, the target sequences of the Affymetrix probe sets which identified the listed genes are also provided in Tables 1 and 2.

Without wishing to be bound by theoretical considerations, it is contemplated that the specific sequences used to detect target genes in the Examples may define specific splice variants or sequences in homologous genes. Therefore in one embodiment, a listed gene for use in the method of the invention is defined by the specific sequence used in said Examples. In another embodiment, a gene for use in the method of the invention is not limited by the specific sequence used in these Examples. Indeed the fact that some genes in Tables 1 and 2 have been identified using different sequences (gene “redundancy”) and confirmatory RT-PCR studies (see

Example 4) provides evidence that usefulness in the method of the invention is not generally limited to the specific sequences used to measure the target gene.

Note, in the event of a discrepancy in the sequence between Tables 1 and 2 and the Sequence Listing, the sequence as provided in the Tables is preferred.

Table 1: as described in priority application US60/619027 filed on 18/10/2004.

Gene Symbol	Gene Title	Affymatrix ID	+ up in sensitive	Sequence	LocusLink	RefSeq Transcript ID	SEQ ID NO.
ACOX2	"acyl-Coenzyme A oxidase 2, branched chain / acyl-Coenzyme A oxidase 2, branched chain"	205364_at		Gtgcagcaattitacagaccctgaccccaatccggagctgaccacgacagcagcgttggaa accagacgtctatcacactccaggctgctaaggctgactctatgctacgtg aagggtttacagaagctctggagaaactgaaaaatgaaccggcaggttcagcaggt gctcaagcgcctctgctgacctcatgccaatacaggaatcttgcataactcgggtgact ttctccatgacgcctccctgctgctggctcccaatggacatggcaagaacgctacccg gacctgctccgcctgatccggaaagatgccatccctgtaaacgaltcttttgactccacc gatcgttttaaatcaagccctggctgttatgtagaacgctctacgaacgcctgttcc agtgggctcagaagctc	8309	NM_003500	SEQ ID NO:1
ACTR2	ARP2 actin-related protein 2 homolog (Yeast)	200729_s_at		gagctaaagctcgtgttttgtaatgctctgtttatccagaagcattaaaggtiaaccocat tgcacagctatcattctgcaaatattctttataataactgaccagtgcttaataaacaag caggctactcaaaataactgctgcagagtggttataatgggtttaaaaataaacatg gaatacagcactgtgccaattgggtlaatttcatlgtgtttgtttgtatttgtaaac ctggaaatacagaataaattgactgttaaaaatgttggccaaaaataaactgaatttaatt tttttaattgactgaaaacctatcaataactgttaattctcagccaactcttgaagctgaaa gaagactcttgattttglaaacggttagcagacattctcgcagctgtagaagaataccta tttatgaatcctcgtggctattctctgtatctgaaaaaaaataccaaatagaccataacatg agttattcttaa	10097	NM_005722	SEQ ID NO:2
APOL1	"apolipoprotein L1 / apolipoprotein L1, 1"	209546_s_at	+	agaatagagaggaggtgaaggaacacagcaalgagaaggccaggaagaagaa agcgtgaaatggagaaagcccaagaggttagaacagtggtacagaggaaga aacagcggctccadacagaccacgcccaggttcaatgtctcccgaaagaatgaag tcttctcctggtgagtgccctgcctctcttcacgataccactcctcctgtctccctgg ggccatatactcagtcagcagcggctcctgatgctgctggtgggtggtgtaatggtg atgggtccctccaggttactaaaagggtgcatgtccctgcttgaacactgaaggcag gtgtgt	8542	NM_003661 / NM_145343 / NM_145344	SEQ ID NO:3
C10orf10	chromosome 10 open reading frame 10 / chromosome 10 open reading frame 10	209183_s_at		aactcactgcctctgctggtgcaattgggagagttcccccattgatgagggccaagata gaatctgaccactcagctaccatccacccctacacccactccacacaggggc ctcatgccaatgctcagggcccacgtgtatgtagagagcagggccactgtccagctgic cacctggggaagctcaagatgctctaaaggccacggcagggcacttggagctctgaaagg acctatgtccatgagccatctgcgcagaagaagggtgagggcatctggagctctgaaagg atcagggctgggacdgatcagaggtgaaagggacacagagagagagaggaggaaga ttgagctggggccaacagccaagctaccctggcaggtctctgccacctctgctct gtgacgtgcagctagghattctcttttttggcctatttttaattgctttgatttgtaaatg ttttctctctctgttaacgtcttt	11067	NM_007021	SEQ ID NO:4

CALM1	"calmodulin 1 (phosphorylase kinase, delta)"	211984_at		tgaaaggcgtgcagtagggccagcgctgctgctgcagaggaatagcttaaca cgaacccctcagcactacgggaatctctctcgaacaacgaatglaaatggtaagt ctactctcogctcaatcaatatttaagcaattgaaatattatglatatcaatattct cccttgagcagtagctagctccactaagctgcttaattctatccctccagctgcagttac tgattttaacccctgaagttgctcgtgagggagacagaatcttctgctgctglatccct ggagtaa	801	NM_006888	SEQ ID NO:5
CALM1	"calmodulin 1 (phosphorylase kinase, delta)"	211985_s _at		ggaggaatggatcctgatalttbagatggcctttgagcactggtgcoagggaagc ttttctgattttgacaaatgaattttgcacacttcaatggctgcttcggcaacttacaca cattgaaat	801	NM_006888	SEQ ID NO:6
CD44	CD44 antigen (homing function and Indian blood group system)	210916_s _at		caagtttggggcagcagcctggggacotcgtccctgctgctgcagcctggcga gatcattgaaataaaccctgcgccttgcaggtgattccacgtggagaaaaatgctc gtacagcatctctggcgggagcgcgtgacotcgtcaaggcttcaataagcact gcccacaatggccagatggagaagcctcgtgacatcgattgagacctgacgtt gcattgcaagcaacagtcgaagaggtggtggcagaagaaaaagctgctgataca cagtggaagggagctgtggaggacagaagcgaagtggaactcaacggagagggc cagcaagctcaggaatggctgattgggaacaaggagctgcagaaacccag accagt	960	NM_000610	SEQ ID NO:7
CD44	CD44 antigen (homing function and Indian blood group system)	212063_at		attgtaaatcttttgcctcagagactcccttaaaattagctcgtgagtaaaatcaa aagagacaaaagacatctcgaatccatattcaagcctggtagaattgctttctag cagaaccttccaaaagtttataatgagatcatalaacaacacaaagaattgattgtag ccaaactcattcaatactgtatatacagaggagtaggagaggaacattgactta ctggaaaaagcaaaatgtaactaagaataaataacatggctcattccacttattgta tagatatgctttgttaaatcaattgtttgagtttcaagaatagccatgttcaattctgt gctgtaacaatgaccactgtattgtactttgactttcagagacacccc	960	NM_000610	SEQ ID NO:8
CDS2	CDP- diacylglycerol synthase (phosphatidate cytidyltransferase) e) 2	212864_at		ttctatgcatccacacacaaacccctgcaagaatgaaatgaaatgcttcttataagag gttccactcgcagacgtgaaggtttcagtttttttttttttttttttttttttttttt tatttataatgctcggagaaaaaacacacatgaaatattcaatgaaatgaaatgaaat tactgaaatgagcccttccacatgtaggtccaaatgctcctgaggtccacctaag ctgtgttttcagggaacatgcaatcattgtgtgtgagactgctgctgctgaatcct ttcggggactttctcagggcagggagcagagggctcctcgtcagcacccttggcc tgaacacacatgtagctgctgtgtgtgtatataatccttaagaggaggtgtgtgtgt gtttgttttaaaagtcacttattctacagtgatttcaattgcacacatgactcttcadtaa accacaaagctcgttataaacctatggaaaaacctaaccctgattagagccttgac	8760	NM_003818	SEQ ID NO:9

[illegible]

DPYSL 3	dihydropyrimidina se-like 3 / dihydropyrimidina se-like 3	201431_s _at		tgaggccacagggtggtagtggaaagggtgttgggaaatgttaaatcagttac ccgtagtagagctattcttgaacttaagtctcgaagtgaagtggaagattgtagtcatcct gaaatgtgttacttcaaatccctcagcctgttctcagcagctgtctatactgagagt gtcatgttccacaaagggtgacacctgagcctgattttcacatccctcaggaagcc ccttcagtgagggtggcaattccaacttccgcaagcttcccaagctccaggtcttcc ctggaaadccagctgagtcacagatacacatcaggtgcccctgggcagccag catcattgtaagtccctcttgaaaacgtgtgtgggtgttcagctgtgtgtgtgggt atggacagacagtaatactctgtgaltgtgtcagctgtgagcgagctctggaaagtg a	1809	NM_001387	SEQ ID NO:15
DUSP4	dual specificity phosphatase 4 / dual specificity phosphatase 4	204015_s _at		ggctccacgcaagggtgtagaacgggtgcccgaatgttccctggggaagttgggactgagc agctgggagcagcgaccgagctcttcccatcttctcctgtgccaacgacgag gccagccagatggcaataagactccgaatacaataaaagcaaacagaaaca ctcaacttagcaataacgggtccgcaagcagcggaagaccttggtgtgtt atgtcagtttcccttccgatagaatacttctacatcttttaagcagtaaggctga agtgataaacccacagatcctagcaaatgtgccaaacacagctttactaaaggggg aggaaagggaaggcaagggtgtagaagacaaagtcccaagtgccctggtctg g	1846	NM_001394 / NM_057158	SEQ ID NO:16
EIF3S4	"eukaryotic translation initiation factor 3, subunit 4 delta, 44kDa / eukaryotic translation initiation factor 3, subunit 4 delta, 44kDa"	208887_at		gatacgtggggcccatgcaagaagagctggccgagcagctgggctgtctactgg cgagaaggagaagctgccgggagagctagagccgtgacggccacgcagaaac aagacagggaagtagtgccgcgagcgtgcgcgagggggccagccgcgcgg ggagtcactcagcccaaacgcagacgcagcagacacacccacccatccgtgcac caactgtcaggggacacgcgtgagacccgtggaagcgtctcctcggcttccgg ctccatccgcctactcctgcttaaggacacacccatggccaatccaaagggttt ggcttcacagcttccacccgcgcgaggtgctgcgtgcccattgccgggtgtccg gcttggctacgacacacatccatccaaagctcagtgaggccaaagccgtcccaact aagccagctgcaactgtgtactcgggtccggaccctggcgacagaagacagcc c	8666	NM_003755	SEQ ID NO:17
EIF5A	eukaryotic translation initiation factor 5A	213757_at		atgtgcggggagagagcccgcaagggaagggaagcccaannggggcagggcc ctccagatgcttgaggagggggaggtccctccctctctctctctctctcccat ciaaagggttggggagagacacagcaggcgagggggtgtgtccacagctgtt gggtgtgtcagggtaagggtatngcaacagggggaacagaccagggatg agtggggaggggcacaaaggaccattgccaagaatccaccg c	1984	NM_001970	SEQ ID NO:18
FADS2	fatty acid desaturase 2 / fatty acid desaturase 2	202218_s _at		ctgtgtccaggaatgcattctgtagaggggggcgaggggtgggctgtgaca atctgcttccacacatggcctgtcctgtgtgtgcccgtgacgtgcaggaggccag gagccagcgggagggagctcagaggaggaggtgtccctgaggggtgtggggag gggtacctcatgaggaacaggtggagctggaagaggaaggaggtgggggt ggaggtgctgtgagctgaggggacggcaggtggaagggggagggaggaagtc ctgggaggtatctgagctgtgtgagcttaacccactaatcagttctagattcagg gaggggacggcccaacacacagaaatgggggtttccggggagggcgtcagtc cccagctciaagcagcaggaaggacatgcatcgaatcgtgggtgtccatggc aatgtgcatgccccacagctatgtgccccgaaccccgacagcagaggcagaatgaa cccataggagagctgaltgtaat	9415	NM_004265	SEQ ID NO:19



FLJ12442	hypothetical protein FLJ12442	218051_s_at			gggaccaccctctatagtgatcgtggcgagatcctatgctggcgccacggctggcgccacag gcgcacatcaccggagctggagcgtgagatccgcacatcaacacacggagcagttac atgcactcgtgacgtggcgagcggcgctcacgggggctgctggcgagcgcgcatgcaga cctatcaggacggcgagtcgagggcaggtgctggctggctggatgaaagagcgga ggagctggatgcacacaaagccctgtcctcaatggcaggtcgccgacgcatctccg cacctccacacccacactactcgaagcgctggctggcttctgacctctacat ggcctccctcagctgctgcacactacccgctggacitcacctctaccacgcogta cgccgctgacgacagggcacccctctggatggaccagctctgcaccggctgcatg aagaccctctcctgtgacatggccacatccgctgaggcgacccttattgtctggg ac	64943	NM_022908	SEQ ID NO:20
FLJ22028	hypothetical protein FLJ22028	213878_at	+		tatitcaaacggagctcccatccaagaaactgaaacccctagtattgttaaaggg ccagctaaatcttcacttacctttacagaaactatattctctctccatccacg aaatcaatcagaaacactgactttctcatgttcaactggacctagggggaattatgacag aaaagcatccataggtcttataatacttttaaaatataaaactgaaatataagc cattccctgaaagagttctggcgactttgtcacttgcatagttaatagcattgctc attgtcagaagattagctttagctctatttcaatacggaaatgtagcataagctgta aaactgtgctctctgcaaaaaaaggcccaacaataagaagaagctttgaaaggga atcaaggaaaaaattataaaagaataactatattgcggaagtaattcttaacacatt gac	79912	NM_024854	SEQ ID NO:21
FNTB	"amesyltransferase, CAAX box, beta"	204764_at			gcaagtcgcgtgattctaccacactgctactgctgagcgccctgtccatagccca gcactcggcgagcgagcctgtgcatgattggtccctgggtgtgcgcgaaacgcct ctgcagcccaactccacgtgtaacaactggacacagacaagggtgatccaggccac tacatacttcacagaagccagctccaggttttgagagcttaaggatgagacatcg gcaagcctgcaacggcactagaggaacctgggtcccgagctctgtctaccaccatc tcccacgtaagcaaggtttatagcttcaatacatalcgtactctgtctacacaagcc ttagccctcgtggagctgtggtctctgtgactctctgtcaacaaacccaalggctctg ggttggagaacacagctggcgtgtttaaattcttccacactgtcaa	2342	NM_002028	SEQ ID NO:22
GPC5B	"G protein-coupled receptor, family C, group 5, member B / G protein-coupled receptor, family C, group 5, member B"	203632_s_at			tgatgtcaactagcagggtctcagggttccactaggatgcagagatgacctctgc tgcctcaacgactgacacctgggttccttccgtgctatgtgaaattctctggagtg gaatggatcacaagaggttctgtcttttgaggggtgggggatalattttgtttgttt tctgaggttccatgaaaacagccctttccaaagcccatgtttctgtcaltggttccatc gtctgagcaagtcattctgtttatagcatttgcacatctcggccactcaaaagcc aatgtctctgcactgttggccagcatalaactctagcactcgtatcaagagcaggttta acctgacggcatggaaatglaaaatgaggttgggtctctctgcaatactctaatcact acaattgtttctataaaactaccataaagccttaaccttaaaagaaaaatgaaaaag gttagtgttggggggcgggggaggactgacgcgttcaataagccaatgactctgagct gaga	51704	NM_016235	SEQ ID NO:23

[illegible]

Gene	Protein	Accession	Feature	Position	Score	Seq ID
KIAA0931	protein	213407_at	+	213407_at	23035	SEQ ID NO:29
KLHL7	kelch-like 7 (Drosophila)	220239_at		220239_at	55975	SEQ ID NO:30
LAMC2	"laminin, gamma 2 / laminin, gamma 2"	202267_at		202267_at	3918	SEQ ID NO:31
MLLT3	"myeloid/lymphoid or mixed-lineage leukemia (trithorax homolog, Drosophila); translocated to, 3" / myeloid/lymphoid or mixed-lineage leukemia (trithorax homolog, Drosophila); translocated to, 3"	204918_s_at	+	204918_s_at	4300	SEQ ID NO:32
MNAT1	menage a trois 1 (CAK assembly factor) / menage a trois 1 (CAK assembly factor)	203565_s_at		203565_s_at	4331	SEQ ID NO:33

[illegible]

[illegible]

PEX3	peroxisomal biogenesis factor 3	203972_s_at		<p>tggaatccaaacccattatgcaattatgatgccagatgaagaaccccaattgacagtgcaggccgtggadttctclogagacattaccattataacdtctcaatgaacctagagacatgttgaaagcccaattttagacgtttgaatacctgtttaaacagggttttagtagacttagaacaatgtgagttcttcgaactactgaacaggacatggaacaatggaacttagaatactcttcacgtgcacgctgccttagtaagaataattcaataglaaacggacacatccattcagttgcagtgaaacacactagtcatttgttcagatctgtgacaatggagcaagacattgtgctaaatgtagtgaagcttttagtaacccctcagcaactcggagaat</p>	8504	NM_000630	SEQ ID NO:42
PIN1	protein (peptidyl-prolyl cis/trans isomerase) NIMA-interacting 1 / protein (peptidyl-prolyl cis/trans isomerase) NIMA-interacting 1	202927_at		<p>agcatttgaaagccctgttgcgtgcggacggggagatgacgggcccgtgttcacggatccggatccacatcatctccgcgacigagtgaggtgggagccaggtcctggcgtggcgagggcggtgagcgccgacgtcccccctgcgcgcagccagtgccgaacccccccatccctgccacggtcacacagattatttgttccacaatggctggagggggccctccagattggggcccctgggtcccccactccctgiccatccacgtggggctgcgcgcgcagatctccctaaaggaattgacttcagcaggggtggagagcctccagaccocaggcaggtgtgtgggaggggtgtccaaaagagaaaggcgtgcagcagccgccctgtcccccaggtgctggaggcagactcgaggccgaattgtttcctagtgagccacgctcctgttcaagtcgcaagggtgaacacatcatgcggcagcgc</p>	5300	NM_006221	SEQ ID NO:43
PRKCA	"protein kinase C, alpha"	213093_at		<p>gattaacgactgtcttggtaacctctgtttaaccttaggagatccaattcctgtgattgtagaacttggatattctctggaaagaataatcattcttctgaagggtgtgttactagaattatcaaaaataatcatgaaggcagtactattttagtctaaagggtttctaaaaaftaacatccatccctctgttagggcttctgaataatctttataaacagaagcatttgaagtcattgttctcaatgattgtgtgtggaaggacatccacgtttaatcattaattgaaatacatataaagcccccactgttgtggaggaagacggaggtgtgtgtttcctctgtatagaacacclactgacaaaatgtagagcccatcaaccgtcaaacacccatttggttatctgcagaggagcaggtgtgtgtaattactgattgcttttttcaagttgtataaactctccgtttgcattgataccgctgttgtagaa</p>	5578	NM_002737	SEQ ID NO:44
RIOK3	RIO kinase 3 (yeast)	202129_s_at		<p>tgaatgtacgctgtccatgctgacctcagtgatataacatgctgtggcatgctggaaaggctgtgtgtgatcgtgcagtgacgtagaaccataccccaacctcaccggtcggagtctgtccgggactgcaggaaatgctcgcagttttccagaagaggaggtcaaggaaagcccttagtgaacggagacactcctcaatgctgttcagggtctaaacatcacagagataatgaagctgatttttagctgagatagaagcttggagaataatgaatgaagatacagttcagaagaatggaaagacgtgctcctatttttgaagatgatggagaccaccactactatagatgaatgacataaacccactgttcaagtgtaaacacagcaggtgattgtagcctgccaatgacaaatgaaatgatgggtgggtgaataataccaaacccactggaatgtgtctgtg</p>	8780	NM_003831 / NM_145906	SEQ ID NO:45

SERP1 NB9	"serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9 / serine (or cysteine) proteinase inhibitor, clade B (ovalbumin), member 9"	209723_at	+	ttgcaccattggccgtgtgtgtgaacctggccctcaagcaatccgccctacacag ccctcccaaaagctcgtatgattacaggaataagccactgagcccaagccctgctcagta tcctttatgtaattataaacatcagcaacattatgatacatatgcagatactatgcatctct ttattagtggtgaagtgctcattgcaatttattgctctgaattccctatcatatgaattgca ttcaacacacactctctgctcgtctttacataatgcttgccttataaagataattatccctct gtttatattctctcattctctgattgcttttaa	5272	NM_004155	SEQ ID NO: 46
SIX1	sine oculis homeobox homolog 1 (Drosophila) / sine oculis homeobox homolog 1 (Drosophila)	205817_at		ccggaggcaaaagagacggccggaggaggaaggaagggagagacaaccca aaacaataactctctccacaacagcagaaccaactctctctctggaagggggca agccgtcatgtccagctcagaaggaattatccctccccaaggtccagaccag aactcgtctctctgctcagggaatatggccacgcagagctcaactattctc tcocgggttaacagcctcgaacccctcaccctcagctctgggacacccacagcaltcag ctcaagadctctcctcggccctcaccctcagctctgggacacccacagcaltcag ggggaggggacgtggggccctcgaagggtattctcggagcaacacacacacacac aggacacacgttaaatagaaatcaggaaacattttgcagctgttctggaggtgtgcg cataaaggaaatgggtggaacttcacaaatatcttttaaaaaatcaaaacacacagcat ctcaagcttaa	6495	NM_005982	SEQ ID NO: 47
SLCO3 A1	"solute carrier organic anion transporter family, member 3A1 / solute carrier organic anion transporter family, member 3A1"	219229_at		ggctgagccaccagtgagctcttggctctactctgacccctagacaacctggggaggga ccctgcccgaacacagacacacataaggcaaaagttatctataactgggaagcc atgagtggtgaaacacatgagctcgtttttatagctaaaggaggggtgaacct gtattgtaattccaaagggtcattttttcttaaaaaaagaaaaaaggttccaaaaaa accaaaactcagtaacacacacacacaggaacagatgcacacacacacagacagac acacagacttgcctttttctcagcatcagagccacagaggtattcagaataagga gaatgacatcgtcggcagggctcggagggccactcgcgcgtgggcccacagag tctacttgaaggccacctcatgttttcaggatgctgacagctgcaagcaacaggcact gccaaattcagggaacagtggtggccagctggaggtggac	28232	NM_013272	SEQ ID NO: 48
SPINK 1	"serine protease inhibitor, Kazal type 1 / serine protease inhibitor, Kazal type 1"	206239_s _at	+	gagacgtgtaagtgccgtgcagtttcaactcagccctggagcgcagaactcagcca tgagggtaacacaggaactctctcagtgccctgtgctcctgtgactctatcgttaacct ggagctgactccctgggaagagagcccaaatgtaacatgaacttaattggtatgcac caagataatgacccctgctcgtgggacgtggaatactatcccaatgaatgcgtgt atgtttgaaggctcggaaacgccagactctatctatcattcaaaaaatcgtggccctgctg agaacccaagggttgaaatcccatcagggtcacccg	6690	NM_003122	SEQ ID NO: 49

SPINK 5	"serine protease inhibitor, Kazal type 5 / serine protease inhibitor, Kazal type 5"	205185_at	+	agccatccatglttagagctctcaagaggaagacagccagacitcttcaagtctctg gattctgagatgtgcaagactaccgagatgtgccaggataggctatctttgccaaa ggatttaagcctgtctgtgtgagatgtgccaaactatacaaatccttgaatgctct gtcatgaaaacctgatacccaaaaataacacacatccgcatgacaggaggtgt gaggagagcaacccagccacccagccagccagcatgcccgtctgacga atgacaggaagatgtgaaagccatgagggaataaataaaccacagttctgaatc acctactccactctgtatatacaagaattctcgagctgtcttattgtctatagaa aacaatacagagctttgggaatggaatcactgatttcaagcttttccatttctctcta gaatctgtgactctgagggtataaagaacattccaccaagtgtgagcccaaaaatgccc tgattacaatgtctgtctgccc	11005	NM_006846	SEQ ID NO:50
STC2	stanniocalcin 2	203438_at		gtccaatctctgaagcatgtatgacatttgacacatcaataaataaagcaaatg agtcatlaaaaataacccctctactgtgctttatctgatacaaaaattactcatgagcc ttctttgaggaaagtgtgacttccaaataaagaattgtgtttttttgagctctgcatc taacaagatgactgaacacactctctgtgtatcaataatagccctgttattctgaagt gagaggaaccaatgtagtaaatgtgacatcaataaactaaataaataaataaataa ccaggccagaaciatagtcatactcacacaaaggagaaatttaaacctcgaaccaa gcaaaaggcttcacgggaataagcatggaataaacaalgtctcaggtgcccactctta aggaggacaacccctgtctcagatggaattggcaccacgtgagctgtctaaagtgtat aatatctgttctactacggatttaggaacaggaactgtacattgtccattgcat tgggggacttattgtgggacttaataaagattcttgaatcaccgggaatatataatg tacagagatcattgatacatgttgaaaggaagcaagtgaaaggtcagagatgaa gtagcgaagtgttggaatatctgtgaaaggatactagttgtgaaatggaagagaca agtattagaccocaaaaagcaaaacagcagagatgcaagagatgcccacaaa ggacaaagcaacaattttctgtgacacttaacccggaagactgtgtgtagaagaa aagaaggcttgtgcaactgtgtggagagagagggggcagggaatgagatgtct gagcgtacagcagacaaagcgtgagcgtgtgtgctctccatccactatgaaatgac taatttaccaggagacccatgtgttatgttctctaatcttccactctccctaaagccctct gagagagatg	8614	NM_003714	SEQ ID NO:51
TAZ	transcriptional co- activator with PDZ-binding motif (TAZ)	202132_at		gocctgtcgcacagatcgaggatgcatctccgggacgttggaacacacagacatga agataaagaaccgtgtacggagctgtatctccaaactgaaggatgccaagaacccgtg accgtgcgggaatgtgtgtggggccataacaccccacgagatcgtgtgtaga ccacagagagatggcagtgatgagctgaaggatccgtaaggccatgaccaa ggaggccatccgaagacacagatggcccgactggcggaacgacagacaccc gttccactcggcgaatgtgagggaataaagaactgcaactacacacaggtgacagcc cgacgtctgtagagccatgaccacctttgtgtctgcaacgaggtgtggaacccgt ggaggtctgtgacccctcgtgtgtagatgtgtgagcctgtgggcccctcccgccca cgctccgtgtgacacagctctctctggagacccatagaaaggcggtgcatgtccc	25937	NM_015472	SEQ ID NO:52
TCEA2	"transcription elongation factor A (SII), 2 / transcription elongation factor A (SII), 2"	203919_at			6919	NM_003195 / NM_198723	SEQ ID NO:53



Gene	Protein	Accession	Length	Seq ID
TNNC1	"troponin C, slow / troponin C, slow"	209904_at	+	7134
TRPM2	"transient receptor potential cation channel, subfamily M, member 2"	205708_s_at	+	7226
VAMP4	vesicle-associated membrane protein 4	213480_at	+	8674
ZNF313	zinc finger protein 313	200868_s_at	+	55905

Table 2: as described in priority application US60/619027 filed on 18/10/2004.

Gene Symbol	Gene Title	Affymetrix ID	+ up in sensitive	Sequence	Locus Link	RefSeq Transcript ID	SEQ ID NO.
AG:CT2L1	alanine-glyoxylate aminotransferase 2-like 1 / alanine-glyoxylate aminotransferase 2-like 1	221008_s_at	+	gctgaagaagccacatagactgcttagggacagcaccactgactccaaa gaaaatccacagcagaagaagaatggaatggtgacggatcacacattactgct cagtaagaggctcaagacatgactgattgcatttaaagcaagatgogctgccc aggttacagagaatgagtagtgcctcctcagcgttaaatagctctattactct aaaagggaatgicagtttagattcaataaagaaaaggtaaaagaglaaacaga ataaaccagatataaicaaacccatgtcaagattattagtcagactagcctggt aatcttctagtgatttcgaagctaccctgatttactattataaatiagactgcaaa ctcaaatataatggcaattactctcatgttttaattggtcaaatagagagca aagtaaacagggtccctcaccttttgagact gtgccatagtgcaggctggggaacttaagcctcactgattataaocccagaaa acagagcctctagatgaacattctgatcaagggtacaaatctttaaaatcacta atgattgagggtccattattagtgactcagaatggtcactttctattacacgga gtgtgtaaaactaaagcatttgaaacatacagaatgtctattgtcatggga aatcttctaaaccagtgagggttagaagaaagtattatcttgtagcaaatia actttacatcttttctactgtatgtgttggtggaocgataaagtgtctaatctga ggcaaaagtagaattgtttatgtatgaagaaagaaatgtgtaagttttga ttctactctatgtctggaactgcattcacacatggcatgaataaagtcagggtctta caaatgttatttgatagatactggaatgtgttgccatattgtgccatt gggatgcatgtggccattgticaaagtgcaagacaaagcctctcttgcg acaaacttaaaaatccatgaagggtgctgcacagatgagagactctgacc aggatcatggtatcccgagtgagattgacctgctcaacatccgagggaatc attgagaatattgacaagctctccacaaagccattgagggtgacaccccgga gactcctgaaggcctgtgctgctctgtgtggtgagactagggccacagctt tgccgggcactctgccaaagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagactaaaggaagggggaaggggtgggggggggg gttgggtgggctctatctcatgagcttaggaacgctccacccccagggc catcgaggggccagcagcgtgagcgggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcgaagaatgcaaatatataagacacctggagca gtacatgcaaatgtcaagcaaatgatttgcgtgacaggtgtgggaaaaagttcc atgaaacacacacagatgattgacagtagttatcaaaaatgcaatgacagcca tagctgctttatgtcgtgtatccacagctgtgtgttattacagtcacgttta gaagacaatcgcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgacatgctatagcataactgaagataaaatta cagatatacaattgagctactgccaaagtcatagccaataaagatgagctggc ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaacttca attgtcacttttatgctatttctgata	64850	NM_031279	SEQ ID NO:58
AKAP12	A kinase (PRKA) anchor protein (gravin) 12 / A kinase (PRKA) anchor protein (gravin) 12	210517_s_at		gtgccatagtgcaggctggggaacttaagcctcactgattataaocccagaaa acagagcctctagatgaacattctgatcaagggtacaaatctttaaaatcacta atgattgagggtccattattagtgactcagaatggtcactttctattacacgga gtgtgtaaaactaaagcatttgaaacatacagaatgtctattgtcatggga aatcttctaaaccagtgagggttagaagaaagtattatcttgtagcaaatia actttacatcttttctactgtatgtgttggtggaocgataaagtgtctaatctga ggcaaaagtagaattgtttatgtatgaagaaagaaatgtgtaagttttga ttctactctatgtctggaactgcattcacacatggcatgaataaagtcagggtctta caaatgttatttgatagatactggaatgtgttgccatattgtgccatt gggatgcatgtggccattgticaaagtgcaagacaaagcctctcttgcg acaaacttaaaaatccatgaagggtgctgcacagatgagagactctgacc aggatcatggtatcccgagtgagattgacctgctcaacatccgagggaatc attgagaatattgacaagctctccacaaagccattgagggtgacaccccgga gactcctgaaggcctgtgctgctctgtgtggtgagactagggccacagctt tgccgggcactctgccaaagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagactaaaggaagggggaaggggtgggggggggg gttgggtgggctctatctcatgagcttaggaacgctccacccccagggc catcgaggggccagcagcgtgagcgggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcgaagaatgcaaatatataagacacctggagca gtacatgcaaatgtcaagcaaatgatttgcgtgacaggtgtgggaaaaagttcc atgaaacacacacagatgattgacagtagttatcaaaaatgcaatgacagcca tagctgctttatgtcgtgtatccacagctgtgtgttattacagtcacgttta gaagacaatcgcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgacatgctatagcataactgaagataaaatta cagatatacaattgagctactgccaaagtcatagccaataaagatgagctggc ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaacttca attgtcacttttatgctatttctgata	9590	NM_005100 / NM_144497	SEQ ID NO:59
ANXA6	annexin A6 / annexin A6	200982_s_at		gtgccatagtgcaggctggggaacttaagcctcactgattataaocccagaaa acagagcctctagatgaacattctgatcaagggtacaaatctttaaaatcacta atgattgagggtccattattagtgactcagaatggtcactttctattacacgga gtgtgtaaaactaaagcatttgaaacatacagaatgtctattgtcatggga aatcttctaaaccagtgagggttagaagaaagtattatcttgtagcaaatia actttacatcttttctactgtatgtgttggtggaocgataaagtgtctaatctga ggcaaaagtagaattgtttatgtatgaagaaagaaatgtgtaagttttga ttctactctatgtctggaactgcattcacacatggcatgaataaagtcagggtctta caaatgttatttgatagatactggaatgtgttgccatattgtgccatt gggatgcatgtggccattgticaaagtgcaagacaaagcctctcttgcg acaaacttaaaaatccatgaagggtgctgcacagatgagagactctgacc aggatcatggtatcccgagtgagattgacctgctcaacatccgagggaatc attgagaatattgacaagctctccacaaagccattgagggtgacaccccgga gactcctgaaggcctgtgctgctctgtgtggtgagactagggccacagctt tgccgggcactctgccaaagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagactaaaggaagggggaaggggtgggggggggg gttgggtgggctctatctcatgagcttaggaacgctccacccccagggc catcgaggggccagcagcgtgagcgggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcgaagaatgcaaatatataagacacctggagca gtacatgcaaatgtcaagcaaatgatttgcgtgacaggtgtgggaaaaagttcc atgaaacacacacagatgattgacagtagttatcaaaaatgcaatgacagcca tagctgctttatgtcgtgtatccacagctgtgtgttattacagtcacgttta gaagacaatcgcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgacatgctatagcataactgaagataaaatta cagatatacaattgagctactgccaaagtcatagccaataaagatgagctggc ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaacttca attgtcacttttatgctatttctgata	309	NM_001155 / NM_004033	SEQ ID NO:60
AREG	amphiregulin (schwannoma-derived growth factor) / amphiregulin (schwannoma-derived growth factor)	205239_at	+	gtgccatagtgcaggctggggaacttaagcctcactgattataaocccagaaa acagagcctctagatgaacattctgatcaagggtacaaatctttaaaatcacta atgattgagggtccattattagtgactcagaatggtcactttctattacacgga gtgtgtaaaactaaagcatttgaaacatacagaatgtctattgtcatggga aatcttctaaaccagtgagggttagaagaaagtattatcttgtagcaaatia actttacatcttttctactgtatgtgttggtggaocgataaagtgtctaatctga ggcaaaagtagaattgtttatgtatgaagaaagaaatgtgtaagttttga ttctactctatgtctggaactgcattcacacatggcatgaataaagtcagggtctta caaatgttatttgatagatactggaatgtgttgccatattgtgccatt gggatgcatgtggccattgticaaagtgcaagacaaagcctctcttgcg acaaacttaaaaatccatgaagggtgctgcacagatgagagactctgacc aggatcatggtatcccgagtgagattgacctgctcaacatccgagggaatc attgagaatattgacaagctctccacaaagccattgagggtgacaccccgga gactcctgaaggcctgtgctgctctgtgtggtgagactagggccacagctt tgccgggcactctgccaaagaaatggtatcagcaccagcccgccatggccaag cctgattgtccagctccagactaaaggaagggggaaggggtgggggggggg gttgggtgggctctatctcatgagcttaggaacgctccacccccagggc catcgaggggccagcagcgtgagcgggtgaaaaaccgtagccatagatcctgt cc atttcaaaattctgattcagcgaagaatgcaaatatataagacacctggagca gtacatgcaaatgtcaagcaaatgatttgcgtgacaggtgtgggaaaaagttcc atgaaacacacacagatgattgacagtagttatcaaaaatgcaatgacagcca tagctgctttatgtcgtgtatccacagctgtgtgttattacagtcacgttta gaagacaatcgcagaaatataagaggagaaagctgagggaacgaagaaa cttcgacaagagaatggaaatgacatgctatagcataactgaagataaaatta cagatatacaattgagctactgccaaagtcatagccaataaagatgagctggc ctcttccagtggtatcaataagacaatggacccttttgtatgattgttttaacttca attgtcacttttatgctatttctgata	374	NM_001657	SEQ ID NO:61

[illegible]

[illegible]

CDKN2A	"cyclin-dependent kinase inhibitor 2A (melanoma, p16, Inhibits CDK4)"	209644_x_at	+	1029	NM_000077 / NM_058195 / NM_058197	SEQ ID NO:71
CEACAM 7	carcinoembryonic antigen-related cell adhesion molecule 7	206199_at	+	1087	NM_006890	SEQ ID NO:72
CHORDC 1	"cysteine and histidine-rich domain (CHORD)-containing, zinc binding protein 1 / cysteine and histidine-rich domain (CHORD)-containing, zinc binding protein 1"	218566_s_at		26973	NM_012124	SEQ ID NO:73
CLU	"clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J)"	208791_at	+	1191	NM_001831 / NM_203339	SEQ ID NO:74

CLU	"clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J) / clusterin (complement lysis inhibitor, SP-40,40, sulfated glycoprotein 2, testosterone-repressed prostate message 2, apolipoprotein J)"	208792_s_at	+	agcagctgaacgagcagcttaactgggtgtcccggtggcaaacctcacgcaa ggggaagaccagctadactcggggcaccacgggtgctccacactcagct cgacgtctccgtgctcactgaggtggtgaggtctctgactcgtatccat cacgtgacggccctgtagaagctccaggaagaacccdaaattatggagac cgtggcggaagacgcctgcaggataaccgcaaaaagacccgggagaggt gagatggatgtgc	1191	NM_001831 / NM_203339	SEQ ID NO:75
COL4A1	"collagen, type IV, alpha 1"	211980_at		gaagagctgtgtcttaacacatagttttaaagactaggaatgaatgaa acatcgttttcatgttcaactcctaaccacaaatattatgtgccaacacaaac ccaggttcatgaataggtgtctattatagtaacatgtactgtgacttattgttta ttctgttaataattttcagggtttaaacacataacacacacacacacacacac caaaagcaac gcttgaac cgggcatcttcgtgtgttctcatgagctgtcacaagacacacacacacacac catgacccagcagggagggcagatctggaccact	1282	NM_001845	SEQ ID NO:76
COL4A1	"collagen, type IV, alpha 1"	211981_at		tcggctacttttggatgcacacacacacacacacacacacacacacac ccgtggtcccccgtctcgtggtgaggtttgaagtgtccatctcagag tgctacggcgtgggacacgtcaatttactacgcaaacacacacacacacacac caccatagagagggagcagatgtcaagacacacacacacacacacacacac gcaggagagctgac cataatgaagcctgactcagcctaatgtcaacacacacacacacacacacac aacagcaac accgtaaagtgctctatagggaatttggctaaacacacacacacacacacac	1282	NM_001845	SEQ ID NO:77
CTSB	cathepsin B	200838_at		tcacctgtagactgtccgtgggagtagctgtccacacacacacacacacac ccgtgtatccatctccaggaggaagacacacacacacacacacacacacac ggagttccataacacaggtgaaggttcccccacacacacacacacacacacac aagtagcttccacattgtcacagaaatcagag	1508	NM_001908 / NM_147780 / NM_147781 / NM_147782 / NM_147783	SEQ ID NO:78
CTSB	cathepsin B / cathepsin B	200839_s_at		tgtgtgtggagcccttggagaacgcaagctcctccacacacacacacacacac gtttgcaatgtcaac attttcgtcagctcgtatcatgtgggaggaagcagcagcagcagcagcagcagc tgt aggaggt aacacagcttttactgttt	1508	NM_001908 / NM_147780 / NM_147781 / NM_147782 / NM_147783	SEQ ID NO:79

DAB2	"disabled homolog 2, mitogen-responsive phosphoprotein (Drosophila)"	201278_at		ggaacgctccagcttcttccgctgtgagcagagctcgaagggtttat tgcgaataaagtttggttgtttgtttatgtgggttttaattgtgtcttgcaccta tgcaggtctgtgaggtatcaacgacacatcaangtacctgaggggaa gaagggtgtatgtcgaagcclaaacgaaggggccaacgacctia tggatgccaagtagaagagctcgtgtttttatgttaagtgatgat atagcataattcttcatcgtctcctcagtcactataaacaagactgaat agctactaaatgtccaaatcciaaagtgcaaacggaggaactattcagg tagtgaattttgaaagctgactgacgacacacactgtttgtacatact aatcctatgtcagagtggttgggggtctgttcagagcatalaaacctaaaggt atagtagcaagggcaacctctaaagaatactgttcagacacctcagttaca gagaattctaaagaataaaggaagcaacatacctctcttagacacttgg atcacaacactaaggacctttlaagagatagctctctcttctgaagatcaatt ctccaaaggccaagattgtcttctccattctgtagctatgcaaatgaggg aagaaactatcatctctccctttttctgtatctttcagctgtttgtctctg ggctcaagtagtatccaccctttcacaagcaacagactc	1601	NM_001343	SEQ ID NO:80
DAB2	"disabled homolog 2, mitogen-responsive phosphoprotein (Drosophila)"	201280_s_at		gctcactacactatcagtcacacaaatgaattttcacttttlaagatgcatctgg tgcbaacacagatcgaagttgtctcnaaagctatgtctgcaaggctgtgc atgctctgtttaaagggaaggacaggtcttctaaatttgggtgatactttgctact atgggaataactgtaaaaaataatcgtatcccaactctgtctctgtatgactct ctctgcccctttatgacacttgaccacaaatgcctctatgttcacagtcagggc acaaactactctgatacagaaggtcttcaagctttatcatcatactgaat ccctcaccctaaaggagaggtgaaagcaagadgcttgaatgggtatgag ggagattgtgcatacccaagccacctggaagaaatgactcactgcaagagaac tgtggattgtgtctcattcaccctggaataaacaacctatctcagaaggaaca a	1730	NM_006729 / NM_007309	SEQ ID NO:82
EMP1	epithelial membrane protein 1 / epithelial membrane protein 1	201324_at		cacaaataaccaggtgaggttagagagatggccagcaaaaactgtggga agatgaactctgtcatatgattcattatcacatgatatagaaggtctgtagtc aaaaaacatactaacattcagacataccaaaaggaatactcacatttgitaaag aagtgaaactatgactggaataaaccatgattccctcttcttctttttctgac attatgctatgaattgcaactctgtgtgattgtctagactgattgggtctct cgtaaat	2012	NM_001423	SEQ ID NO:83
EMP1	epithelial membrane protein 1	201325_s_at		tatcgccgtgagaagatctacccagggagaatctgagacatctgcactttc ttatagctctccatccactttttatcccttcttttgggaggttatgcaatg attttgttattatgaaaggattattactatcttctctattgtattctagtaag aaactcttgacnnnaagccaccacaaattactaa	2012	NM_001423	SEQ ID NO:84

EMP1	epithelial membrane protein 1	213895_at		aaggacggglatcttctcgtgagcaataaaggagcgggataaagacgtgcatatccttg tgcnnnnnccagcancnalaacaataaaggaggggttttaattggaagcaggcaat ctnccagccctcgtgcttgatgaatagttgcacagagattgacccaana atacacaatggaggctgaagaagttcaacataatttaagccaattaalcaaatgca ttgattctgagcttctctagaaggctctcatgattcttagattgctctgataaacatc ataagggttccacnccctcattttagctcccccagggaatttcttccccatgca tacaccagctcctaaatacaaccccccaaggctatctctccatccctcgcagagg gaactttgcaacatcgcacaaacacccctagctctatccagagctgctctgctg ctaagattgglatcttctcccaaaagccggatggatggatgggggtgaattgagtc agaattcc	2012	NM_001423	SEQ ID NO:85
EREG	epiregulin / epiregulin	205767_at	+	taaaaactglatctgaaccacttgaattttgctccaalatacattctgagactttt gaaaaaaagtttttaattgatgccaaalatactgaccgttaaaaaaatctctgtc alatgggagagggggaggaatgactgtacaacacagatttctgggtatattta atgttttaaaaaagaglaattcaatttaaatatctgtatcaaatgattgattaaat gaaataatgattttcttttttttttgcactcgtgaattgcatttttaagttgaagac caatttggtaaacggtttttaaaagatgctatggaacalaaagttgattgcatgca atttaagtaacttattgactatgaattattatcgattactgaattgatacaattggtt glttcaatatacagcttgataattgtaacctaaag gggactcatttagctctagaccacatacaaaagttcggcatgagctcatgat ctatgctgttctatgctggaagcaccggatgggggtgtagcaaatctgccc tgcacagcagcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactgaaatcactaactgactgaaaatgaaatgggaataagct ttgtccacagatgctgggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgttg atattctcagggctgacagggctcctggaacattccacatatttttaactgca gtataagtcagaaaaataaagttaacataacttccaclaacaacacacatagag attccacaaatccactataattggcaagtggttgaatataatttttagtaatt gcatgcaaaattttctagctccactcttctccctgcttcttttttggggagcig gtaactgataaactcttccocacccttctcaggaataataagtggtttgttgg taacgtgatacatctglatgaalgaacattggagggaacacatctactgaattct gtaatttaaaatattttgctgctgtagtaactatgaacagatagaagaatctacagat gctgdataaataaggaataataaatttccactcaataaataatgctatttttaaat ctattctctattgatttctaatcaagatgattactcttatttcttat	2069	NM_001432	SEQ ID NO:86
FGF2	fibroblast growth factor 2 (basic)	204421_s_at	+	gggactcatttagctctagaccacatacaaaagttcggcatgagctcatgat ctatgctgttctatgctggaagcaccggatgggggtgtagcaaatctgccc tgcacagcagcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactgaaatcactaactgactgaaaatgaaatgggaataagct ttgtccacagatgctgggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgttg atattctcagggctgacagggctcctggaacattccacatatttttaactgca gtataagtcagaaaaataaagttaacataacttccaclaacaacacacatagag attccacaaatccactataattggcaagtggttgaatataatttttagtaatt gcatgcaaaattttctagctccactcttctccctgcttcttttttggggagcig gtaactgataaactcttccocacccttctcaggaataataagtggtttgttgg taacgtgatacatctglatgaalgaacattggagggaacacatctactgaattct gtaatttaaaatattttgctgctgtagtaactatgaacagatagaagaatctacagat gctgdataaataaggaataataaatttccactcaataaataatgctatttttaaat ctattctctattgatttctaatcaagatgattactcttatttcttat	2247	NM_002006	SEQ ID NO:87
FGF2	fibroblast growth factor 2 (basic) / fibroblast growth factor 2 (basic)	204422_s_at	+	gggactcatttagctctagaccacatacaaaagttcggcatgagctcatgat ctatgctgttctatgctggaagcaccggatgggggtgtagcaaatctgccc tgcacagcagcaccatagcagctgactgaaaatcagcactgctgagtagttt gatcgtttaactgaaatcactaactgactgaaaatgaaatgggaataagct ttgtccacagatgctgggggaacccctccactcaagatggaatttctcccca aggattcaagatgaattgaatttttaatacaagatagtgctttattctgttg atattctcagggctgacagggctcctggaacattccacatatttttaactgca gtataagtcagaaaaataaagttaacataacttccaclaacaacacacatagag attccacaaatccactataattggcaagtggttgaatataatttttagtaatt gcatgcaaaattttctagctccactcttctccctgcttcttttttggggagcig gtaactgataaactcttccocacccttctcaggaataataagtggtttgttgg taacgtgatacatctglatgaalgaacattggagggaacacatctactgaattct gtaatttaaaatattttgctgctgtagtaactatgaacagatagaagaatctacagat gctgdataaataaggaataataaatttccactcaataaataatgctatttttaaat ctattctctattgatttctaatcaagatgattactcttatttcttat	2247	NM_002006	SEQ ID NO:88



[illegible]

ITGB2	"integrin, beta 2 (antigen CD18 (p95), lymphocyte function-associated antigen 1; macrophage antigen 1 (mac-1) beta subunit) / integrin, beta 2 (antigen CD18 (p95), lymphocyte function-associated antigen 1; macrophage antigen 1 (mac-1) beta subunit)"	202803_s_at		202803_s_at	3689	NM_000211	SEQ ID NO:94
KHDRBS3	"KH domain containing, RNA binding, signal transduction associated 3 / KH domain containing, RNA binding, signal transduction associated 3"	209781_s_at		209781_s_at	10656	NM_006558	SEQ ID NO:95
KRT13	keratin 13 / keratin 13	207935_s_at	+	207935_s_at	3860	NM_002274 / NM_153490	SEQ ID NO:96
LASS6	LAG1 longevity assurance homolog 6 (S. cerevisiae)	212446_s_at	+	212446_s_at	253782	NM_203463	SEQ ID NO:97

LTBP2	latent transforming growth factor beta binding protein 2 / latent transforming growth factor beta binding protein 2	204682_at	99gagcgaaggccttatacgtctaaagaaatattcaatgtagtgaatcggccca gtgatagcctgtggccacacagcagagggctgccatgggatacagcaccaca tctacaagacccttattacataaacacgcttcttaacaggaaacaaacctctctg ggatcctctttgtgaaacacagtttgatgctctaaagaaagaaagctctattccag tgggtctgtgacaggaagcagcagatctcaatgctgttttccctccactcagaa acccctgccccttccctcagaaacagatggcaggcatctctcagtttacaagc agagactcactcccaaccacactagcggg	4053	NM_000428 / NM_032035	SEQ ID NO:98
MAP4K5	mitogen-activated protein kinase kinase 5	203552_at	acacacatgcaatttgcitaaacaaagttatttataatcacagtttcatcacgaa ttaccitaaaaggagctctatgtttcaaciacagatagttgaaggatontaca gaagattatgatagttgaatattctgaaggggtgtgtatgctagctgtgc taccatgtatgattctgacaaagcagataaaataccgtgattttcttacaatta gggataatgcaataaggaatattctcatatataatccatccatgagcagg ggaaatttaattgccatgataatgattttactatactatgccagagaggaact ataagtaattacacatgtaactgtgggtttcacataatgaggtatcatttgagta gggtgaagaaagaaataatttaaatgaattgaattccatgaggtatgagta aat	11183	NM_006575 / NM_198794	SEQ ID NO:99
MAP4K5	mitogen-activated protein kinase kinase 5	203553_s_at	gaactcgaatctcattgtttacagaaattgtgcaggcagcagcagttagatt coattcagtaacacagttggagagagataccgttttagtgtttgacaaattgt gaaaattgtaaatctacaaggaataattaaatcaagtaagaaactggcctctga gttaagtttgatttcgcaatgaaatctgtagatgactcaggaactgtgtggtcttc tggaaacatgggatgcagggttaaagctcaagctcagatgaggtacocagga gatttgaatgaacaaaggtttccgcttattgatacagacagaggtgtgtgtttg gaaatgagcccaagaaatctctacacacacacacacacacacacacacacac ggacatgaaaatgttactaagcaacagaaactgactcacaatgacagga atgaatatactcattgaaagggaataataaggaaattcaalacaaactgcacta tattgtcttaact	11183	NM_006575 / NM_198794	SEQ ID NO:100
MMP2	"matrix metalloproteinase 2 (gelatinase A, 72kDa gelatinase, 72kDa type IV collagenase) / matrix metalloproteinase 2 (gelatinase A, 72kDa gelatinase, 72kDa type IV collagenase)"	201069_at	ctcagagccaccocctaaagagatcctttgalattttcaacgcagccctgttggg ctgcccgtgtgtgccacactcaggctctctcttcaaacctctgtggtcac agaaacctggagcgaatggagactgtctcaagaggccactgtgtgcccagac agcctggcacagggcaggtggacagggcagtgccaggtggccactccagac ccctggcttttcaactgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgtgt gcaactgtttttttgt aggacacaggt gttccctgttcaacttacttagcatgtccacacaggtgtgtgtgtgtgtgtgtgtgt ggaaacacacagccgt tcccatgggaaat	4313	NM_004530	SEQ ID NO:101

MYC	v-myc myelocytomatosis viral oncogene homolog (avian) / v-myc myelocytomatosis viral oncogene homolog (avian)	202431_s_at	+	<div>gcaacaacgaaatgcacacgccacggctcctcgacacccgagagaaatg tcaagggcgaacacacacacgtcttgagcgcgaagaggaacgagctaa aacggagcttttgcctgcgtgaccagatcccgaggtgaaacaacatga aggcccaagggtatctcttaaaagacacacgatacctctctcgtcc aagcagaggaacaaagctcattctgaagagactgttcggaaacgag agaacgtgaaacacaaactgaacagctacggaacactctgcgtaaggaa aagtaaggaaacgactctcacaagaatgtctgagcaalcacatga ctgttcaaatgcatgaatgaacccacacactgtgctgagctc</div>	4509	NM_002467	SEQ ID NO:102	
NRP1	neuropilin 1	210510_s_at		<div>aacatccgctgtaacacgtcgtctggtgggacctccacccgacactc cctacatcaatgagtgctocaaatagacctgggggaggaagatcgtgag gggcatcattcagggtgggaagcaccggaggaacaaagggtcatgagg aagttcaagatcgggtacacgaacacgctcggactggagatgatcgtg atgacagcaaacgaaaggcgaagcttttggggcaacacacatgatata ccgagctggcgaactttccagctctccacggatcaccagatcccccga gagagccactcatggggactggggtcgaatgtgagctgtgggctgtgag tggaaagccctacacgtggacacacacacccacgggaactgtggatgaa tgtgatgacgacagggccaactgcacagtggaaca</div>	8829	NM_003873	SEQ ID NO:103	
NRP1	neuropilin 1	212298_at		<div>gcaatatctacacggcgcctatgaattacccaaagaaagcttgggtttg gtggttttcatcgcctgtggaatgagattttatgatctctccacatgct agacgctcactcaaaagacatttgtgggagtcacattgcatcatagagac agtcactcatctagttaattgattggaatgcctttgttccaggaaatattg atccactgaagaaagaaatgtttttgcccagagacattatgtgatata atctccacgaaggaaagcactaaagaaacacgtgtgttttaaggcaac agacttaagttgtcctbagccaagg</div>	8829	NM_003873	SEQ ID NO:104	
OLFM1	olfactomedin 1 / olfactomedin 1	205591_at	+	<div>cagggtatctgcacagtgctgcccacacagacacatgttccacggatgc ccgcacaaacagctcggcagctacgtggaaggtgcgaacatctctcaa tccatagagcttgacacggcgaccacagagacttgcagctgtgagaa gatggaacacaaatgaaagactgtgagctcgaaggtcaacacagtgga gagctataagcaacacgtggcagggttaaggggtacatcaaaagagtttt tcaatgtcagtgactgaagaagcagctccactccatgaacacatgaagag agocagagagcttttgcacatgcatcttactatttccatctactagcaact tcaatgaagacatgaatacacaagatcctcttgcacgacgtgagctgc gcgggccacagactcgggaagaaactcctgtttgcagctggaactgagcc caegggcccccgtttctcccccgtctcctctctgttcaacaacacatata aagagggcagagcaatgactgtggcagctcaccggggaacacacac gttggatggcatgaacattcttgatcgtgtgcacgtccggaggaatgtggn nccaggtcttgangagccatggcgtgacccngccgaggtatgtgaa ctcgcacacatgagtgccngtcttactgtgtgtctctctatgataacgt gctgaggtctcnacatagctcctnlgacccgtgctnlgatcatcactgaagcgt ggtcaggtgtgagagtgaaagtgtgtgcccacactgttgtaactcgtaccc cgtgatacactgtgaacgttctctgttactcctgtgaggtgtgaactcgtatgttc aqt</div>	10439	NM_006334 / NM_014279 / NM_058199	SEQ ID NO:105	
OLFM1	olfactomedin 1	213131_at	+		<div>gcgggccacagactcgggaagaaactcctgtttgcagctggaactgagcc caegggcccccgtttctcccccgtctcctctctgttcaacaacacatata aagagggcagagcaatgactgtggcagctcaccggggaacacacac gttggatggcatgaacattcttgatcgtgtgcacgtccggaggaatgtggn nccaggtcttgangagccatggcgtgacccngccgaggtatgtgaa ctcgcacacatgagtgccngtcttactgtgtgtctctctatgataacgt gctgaggtctcnacatagctcctnlgacccgtgctnlgatcatcactgaagcgt ggtcaggtgtgagagtgaaagtgtgtgcccacactgttgtaactcgtaccc cgtgatacactgtgaacgttctctgttactcctgtgaggtgtgaactcgtatgttc aqt</div>	10439	NM_006334 / NM_014279 / NM_058199	SEQ ID NO:106

OSMR	oncostatin M receptor/ oncostatin M receptor	205729_at	+	9180	NM_003999	SEQ ID NO:107
PCDHAC 2	"protocadherin alpha subfamily C, 2/ protocadherin alpha subfamily C, 2"	210674_s_at		56134	NM_018899/ NM_031883	SEQ ID NO:108
PDGFRA	"platelet-derived growth factor receptor, alpha polypeptide / platelet- derived growth factor receptor, alpha polypeptide"	203131_at		5156	NM_006206	SEQ ID NO:109
PDZK1	PDZ domain containing 1 / PDZ domain containing 1	205380_at		5174	NM_002614	SEQ ID NO:110

[illegible]

PLAU	"plasminogen activator, urokinase / plasminogen activator, urokinase"	211688_s_at		5328	NM_002658	SEQ ID NO:115
PP1F	peptidylprolyl isomerase F (cyclophilin F) / peptidylprolyl isomerase F (cyclophilin F)	201489_at		10105	NM_005729	SEQ ID NO:116
PP1F	peptidylprolyl isomerase F (cyclophilin F)	201490_s_at		10105	NM_005729	SEQ ID NO:117
PTGS2	prostaglandin- endoperoxide synthase 2 (prostaglandin G/H synthase and cyclooxygenase) / prostaglandin- endoperoxide synthase 2 (prostaglandin G/H synthase and cyclooxygenase)	204748_at	+	5743	NM_000963	SEQ ID NO:118

RRM1	ribonucleotide reductase M1 polypeptide	201476_s_at	+	6240	NM_001033	SEQ ID NO:119
SEMA3B	"sema domain, immunoglobulin domain (lg), short basic domain, secreted, (semaphorin) 3B / sema domain, immunoglobulin domain (lg), short basic domain, secreted, (semaphorin) 3B"	203071_at	+	7869	NM_004636	SEQ ID NO:120
SERPINE 1	"serine (or cysteine) proteinase inhibitor, clade E (nexin, plasminogen activator inhibitor type 1), member 1"	202627_s_at		5054	NM_000602	SEQ ID NO:121
SERPINE 1	"serine (or cysteine) proteinase inhibitor, clade E (nexin, plasminogen activator inhibitor type 1), member 1"	202628_s_at		5054	NM_000602	SEQ ID NO:122
SLC20A1	"solute carrier family 20 (phosphate transporter), member 1 / solute carrier family 20 (phosphate transporter), member 1"	201920_at		6574	NM_005415	SEQ ID NO:123



SPRY1	"sprouty homolog 1, antagonist of FGF signalling (Drosophila)"	212558_at		10252	NM_005841 / NM_199327	SEQ ID NO:124
SPRY2	sprouty homolog 2 (Drosophila) / sprouty homolog 2 (Drosophila)	204011_at		10253	NM_005842	SEQ ID NO:125
SFIPUL	sushi-repeat protein / sushi-repeat protein	205499_at		27286	NM_014467	SEQ ID NO:126
TCF8	transcription factor 8 (represses interleukin 2 expression) / transcription factor 8 (represses interleukin 2 expression) / transcription factor 8 (represses interleukin 2 expression)	208078_s_at	+	6935	NM_030751	SEQ ID NO:127
TGFA	"transforming growth factor, alpha / transforming growth factor, alpha"	205016_at		7039	NM_003236	SEQ ID NO:128

TGFBP2	"transforming growth factor, beta receptor II (70/80kDa) / transforming growth factor, beta receptor II (70/80kDa)"	208944_at	7048	NM_003242	SEQ ID NO:129
TIEG	TGFB inducible early growth response / TGFB inducible early growth response	202393_s_at	7071	NM_005655	SEQ ID NO:130
TIMP3	"tissue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201147_s_at	7078	NM_000362	SEQ ID NO:131
TIMP3	"tissue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201148_s_at	7078	NM_000362	SEQ ID NO:132
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TIMP3	"issue inhibitor of metalloproteinase 3 (Sorsby fundus dystrophy, pseudoinflammatory)"	201150_s_at		gacatttggaaatgacctgctagaggccaaactgtggccccagaggagacacaccc ctccatgccccagacctgtctgcatgtgacaattgacaattctggactacccc aagatggccaccaggtgttggctctgctacacaaaggttaacatgctacatgag tattttatgagagacaaacattatataaactcagtcgcaaaagcaaaacaaat ggaaagtagggagggtgagatgacacacactccaaatggctctggaggc gagagaaaggaggagactggagaaatgtttgtttggggtagaggtctct agattccacagactccgttccctttagccagctgctgctgaaacccagaa gtgagagagaaacacaaagagatcgaacccctgctagaaaggatglat ttgtgctaaattctgtagccactgtttacagtttccatctgattttatg gaglatctagagcttggccactctccattttgtcccttggctcatttaattggct aatgccccccaaacagaaatatacacaacaaataactaataatgcccacaa aaggcaagactgacctgaaatctagacctgttggagatactaaactgctcga ggaaagtagcttggatgacatgacgaacatgttgcacaaagatgataa aatalgattctattttcccccaccccgaaatgtcaaaatgtccatgtaaac ctgctacaaatggcaggttatatacagaaatgtcaaaatgtccatgtaaac aatgtctgtcctacaccccaagtttcaaagtattaaatgtctctctactactc cgtttaatatcttgaaggttattaaatgtattaaatgtattaaatattattttc tgtaaatgaaactgtgaagatgataaaactgaagcagataactggaaccac ciaaagactccattttgaggtatttttggccctgtgttggattat accaggctctcatgaatccacacaaactgaacacgtggcaataaattia tctgattgactgagtaataatatacaccactattgtgagctgacacactaagtc aagtaaaaggcttggcgaagatgtgtcaatgaagcacaataatgagaga tcaagatgacatgtccaaagacacagacagaaaggttcaactgtctgt aatgtgacacactatggaagaaagagcgtatgacacatgattaaagat ctcaaaagccaactcttctgactctgagagaaatcagacatctatctcaca ggacattactgtgactcagaataatcaactcagaatgaatccaaagctt ggctagagtgaaacacacaaatcagttctgagatatagcaattgattigan aagattcttaagctgctgtaaatctgctgtgtt	7078	NM_000362	SEQ ID NO:134
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Example 4**RT-PCR Confirmation Studies**

In addition, the sequence of the RT-PCR primers used in the confirmatory follow up studies as highlighted in Figs 3, 4, 5 and 6 are listed in Table 3. Note that DAPK2 was not identified by Affymetrix analysis, only via follow up of the DAPK gene family by RT-PCR following discovery of predictivity of DAPK1. Hence no Affymetrix ID or Affymetrix ID sequence is provided for DAPK2.

Table 3

Sequences relevant to genes followed up by RT-PCR (see Figs 3, 4, 5 & 6)

(all sequences written 5'-3')

Gene	affy id	affy probe seq	Tagman Forward Primer	Tagman Reverse Primer	Tagman probe
EMP1	201324_at	CACCAAATTACCTAGGCTGAGGTTAGAGAGATTGGCCAGCAAA AACTGTGGGAAGATGAACCTTTGTCATTATGATTTTCATTATCAC ATGATTATAGAAGGCTGTCTTAGTGCAAAAAACATACTTACATT TCAGACATATCCAAAGGGAATACTCACATTTTGTAAAGAAATT GAACATGACTGGAGTAAACCATGTATTCCTTATCTTTTACTT TTTTCTGTGACATTTATGTCCTCATGTAATTGCATTACTCTG GTGGATTGTTCTAGTACTGTATTGGGCTTCTTCGTTAAT	AGCCATCCTG CCCTTCTGA	ACCTTACAAAC TCTCTTTCC	CAAAGCA AAACATC ACATTCC AGTC
NES	218678_at	GCAGCACTCTTAACCTACGATCTCTTGACATACGGTTTCTGGC TGAGAGGCCCTGGCCCGCTAAGGTGAAAAGGGGTGTGGGCAA AGGAGCCTACTCCAAGAATGGAGGCTGTAGGAATATAACCTC CCACCTGCAAAAGGGAATCTCTTGCTGCTCATTCTCATAGG CTAAGTCAGCTGAATCCCGATAGTACTAGGTCCCTTCCCTCC GCATCCCGTCAGCTGGAAAAGGCTGTGGGCCAGAGGCTTC TCCAAAGGGAGGGTGACATGCTGGCTTTTGTGCCAAGCTCA CCAGCCCTGCGCCACCTCACTGCAAGTAGTGACCATCTCAC TGCAGTAGCACGCCCTCCTGGGCCGTCTGGCCTGTGGCTAAT GGAGGTGACGGCACTCCCATTTGTGCTGACTCCGCCCATCCCT GCCACGCTGTGGCCCTGCCTGGCTAGTCCCTGCCTGAATAAA G	GCCCCCTTCA GGAGGAGGA	AGTGCCGGGG AGATGGTCTT	AGTGCTC TGAAGAC CTCTTGG GC
DAPK1	203139_at	CCTCCTCCAGGGTGATTTTATGATCAGTGTGTTGCTCTAGGA AGACATTTTCCGTTTGCTTTTGTTCGAATGTCAATGGTGAACG TCCACATGAAACCTACACACTGTCATGCTTCATCATTCCTCTC ATCTCAGGTAGAAGGTTGACACAGTTGTAAGGGTTACAGAGAC CTATGTAAGAATTCAGAAGACCCCTGACTCATCATTTGTGGCA GTCCCTTATAATTGGTGCATAGCCAGATGCTTTCCACATTTAG ATCCTGTGTTTCATAACTTCCTGTACTTGAAGTCAAAAGCAGAA AATAAAGGAAGCAAGTTTTCTTCCATGATTTTAAATTGTGATC GAGTTTTAAATTGATAGGAGGGAACATGTCCTAATTCTTCTGT CCTGAGAA	AGGAAACGCT ACCTCTCTGT	CTGGAGGAGG ATTCCCTTCT	CTTGCTG TATGCTG ATCATCG CC
DAPK2	Not applicable	Not applicable	GGGTAGGCAC CTGGCATC	AGTGCAGTGG CGTGATCTC	TACTCCA GGGGCT GAGGTGA CA

### Example 5

#### **Diagnostic test for Clinical Studies**

The predictive gene lists above have been generated using the preclinical studies described. The following approach is employed to develop a diagnostic test for the clinical setting based on this data.

- a) Identify patients which represent the population of individuals whom we would expect to derive benefit from a diagnostic test, and for which pre-treatment tumour samples and outcome of gefitinib treatment are known or will be available. For each sample the expression level for our genes of interest is evaluated, using for example the RNA signal from RT-PCR. QC procedures are applied to identify the set of samples and genes to take forward to step b).
- a) Identify a subset of the genes which together are able to distinguish between patients showing different responses to gefitinib. There are a variety of methods which are useful to select the subset of genes and combine their expression values to provide a prediction, possibly a predictive value and a corresponding threshold which distinguishes between different patient groups. An example is stepwise Linear Discriminant Analysis where genes that distinguish well between patient groups are successively added to a linear combination until addition of a further gene does not provide additional predictive power (Mardia et al.). The threshold value of the linear combination is then selected to give the appropriate sensitivity and specificity properties.
- d) Tool validation would partly be carried out during development in step 2, for example using cross validation and permutation tests. In addition, the finally developed diagnostic procedure (gene subset and method of combining to generate a prediction and a platform for biological analysis) is tested and validated in its entirety using an independent set of samples not used within tool development in step b).

### References

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Kris et al. (2003) JAMA, 290, p2149  
Lynch et al.(2004) New England Journal of Medicine, 350(21) p2129

Mardia K.V., Kent J.T., Bibby J.M. (1979) "Multivariate Analysis" London, Academic Press Inc. Ltd.

Paez et al. (2004) Science, 304 p

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Storey (2003) "Statistical Significance for Genome Wide Studies" PNAS, vol 100, issue 16, pp 9440 – 9445

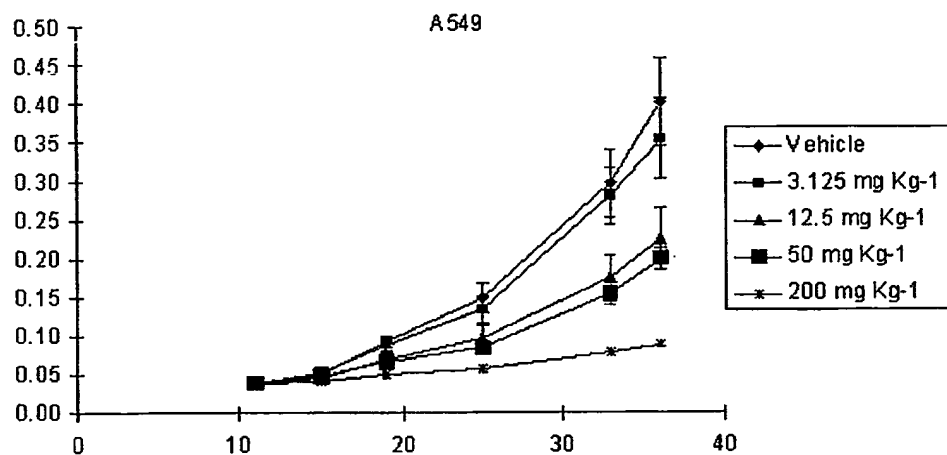
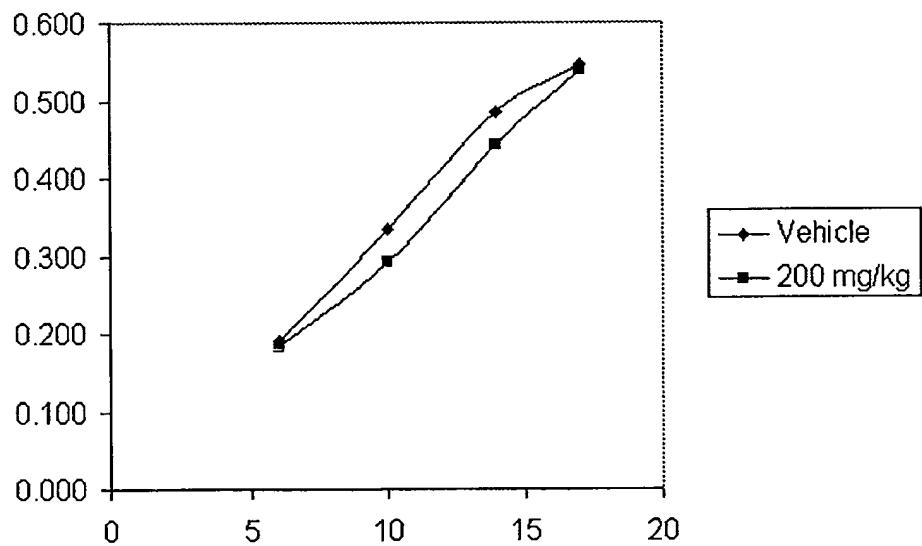
Yarden & Sliwkowski (2001) Nature Reviews Molecular Cell Biology, 2, p127

**CLAIMS**

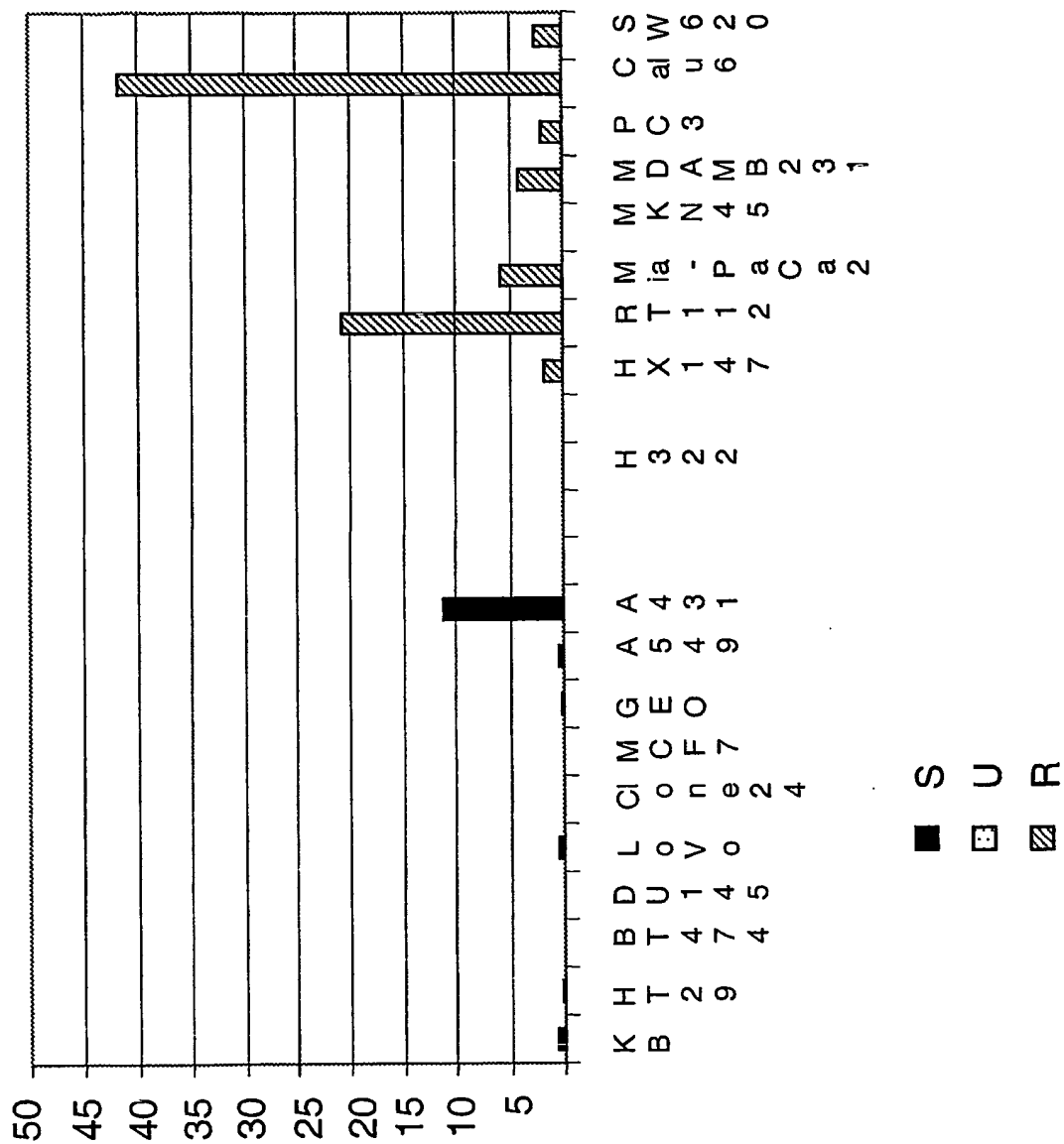
1. A method of selecting a mammal having or suspected of having a tumour for treatment with an erbB receptor drug which comprises testing a biological sample from the mammal for expression of any one of the genes listed in Table 1 or DAPK2, whereby to predict an increased likelihood of response to the erbB receptor drug.
2. A method according to claim 1 comprising testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7, DAPK1, ACOX2, GSPT2, TNNC1 or DAPK2.
3. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of any one of NPAS2, NES, CHST7 or DAPK1.
4. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of at least two of NPAS2, NES, CHST7 or DAPK1.
5. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of at least three of NPAS2, NES, CHST7 or DAPK1.
6. A method according to any preceding claim comprising testing a biological sample from the mammal for expression of NPAS2, NES, CHST7 and DAPK1.
7. A method according to any preceding claims additionally comprising testing a biological sample from the mammal for expression of any gene listed in Table 2 as defined herein.
8. A method according to claim 7 comprising testing a biological sample from the mammal for expression of any one of EMP1, SLC20A1, SPRY2 or PGM1.
9. A method according to any one of claims 7-8 comprising testing a biological sample from the mammal for expression of EMP1.



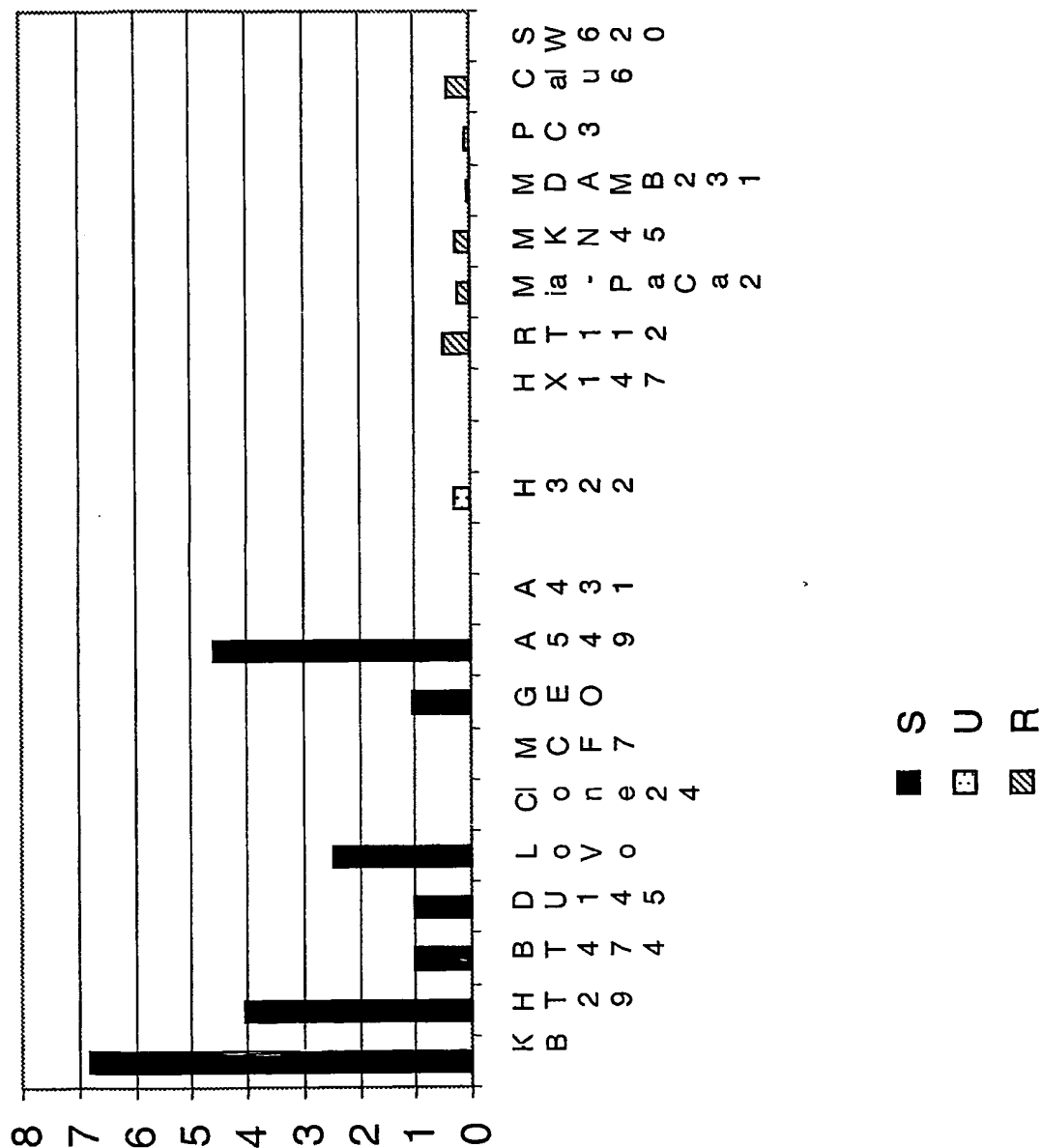
10. A method according to any preceding claim wherein the tumour is selected from the group consisting of leukaemia, multiple myeloma, lymphoma, bile duct, bone, bladder, brain, CNS, glioblastoma, breast, colorectal, cervical, endometrial, gastric, head, neck, hepatic, lung, muscle, neuronal, oesophageal, ovarian, pancreatic, pleural membrane, peritoneal membrane, prostate, renal, skin, testicular, thyroid, uterine and vulval.
11. A method according to claim 10 wherein the tumour is selected from one of non-small cell lung, pancreatic, head or neck.
12. A method according to any preceding claim wherein the erbB receptor drug is selected from any one of gefitinib, erlotinib, PKI-166, EKB-569, HKI-272, lapatinib, canertinib, AEE788, XL647, BMS 5599626, cetuximab, matuzumab, panitumumab, MR1-1, IMC-11F8 or EGFR11.
13. A method according to claim 12 wherein the erbB receptor drug is gefitinib.
14. A method according to any preceding claim wherein the mammal is a human and in which the method comprises testing a biological sample from the human for increased expression of DAPK1 and decreased expression of NPAS2, NES, CHST7 and EMP1 whereby to predict an increased likelihood of response to gefitinib.

**Figure 1****Figure 2**

**Figure 3**



**Figure 4**



**Figure 5**

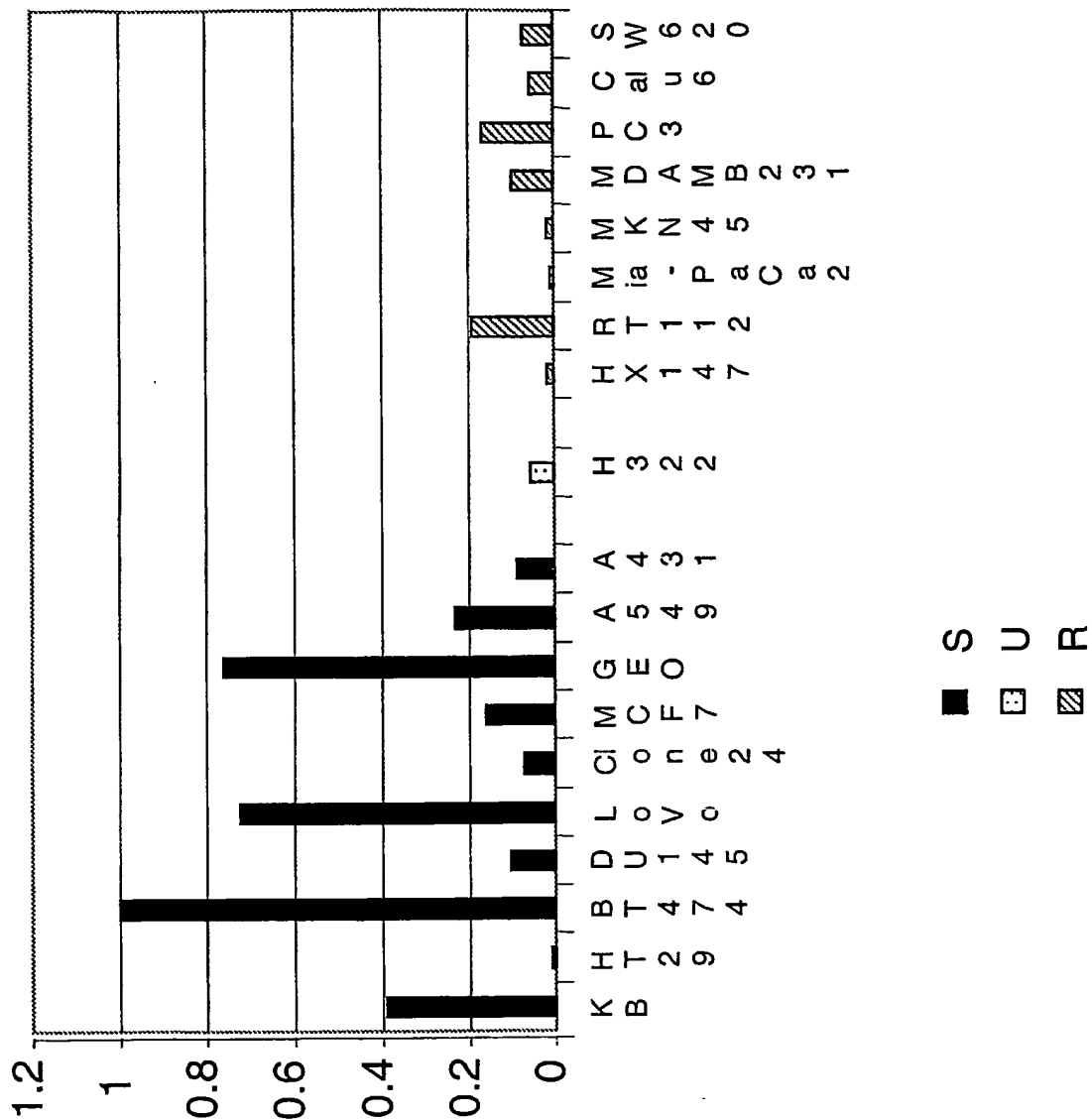
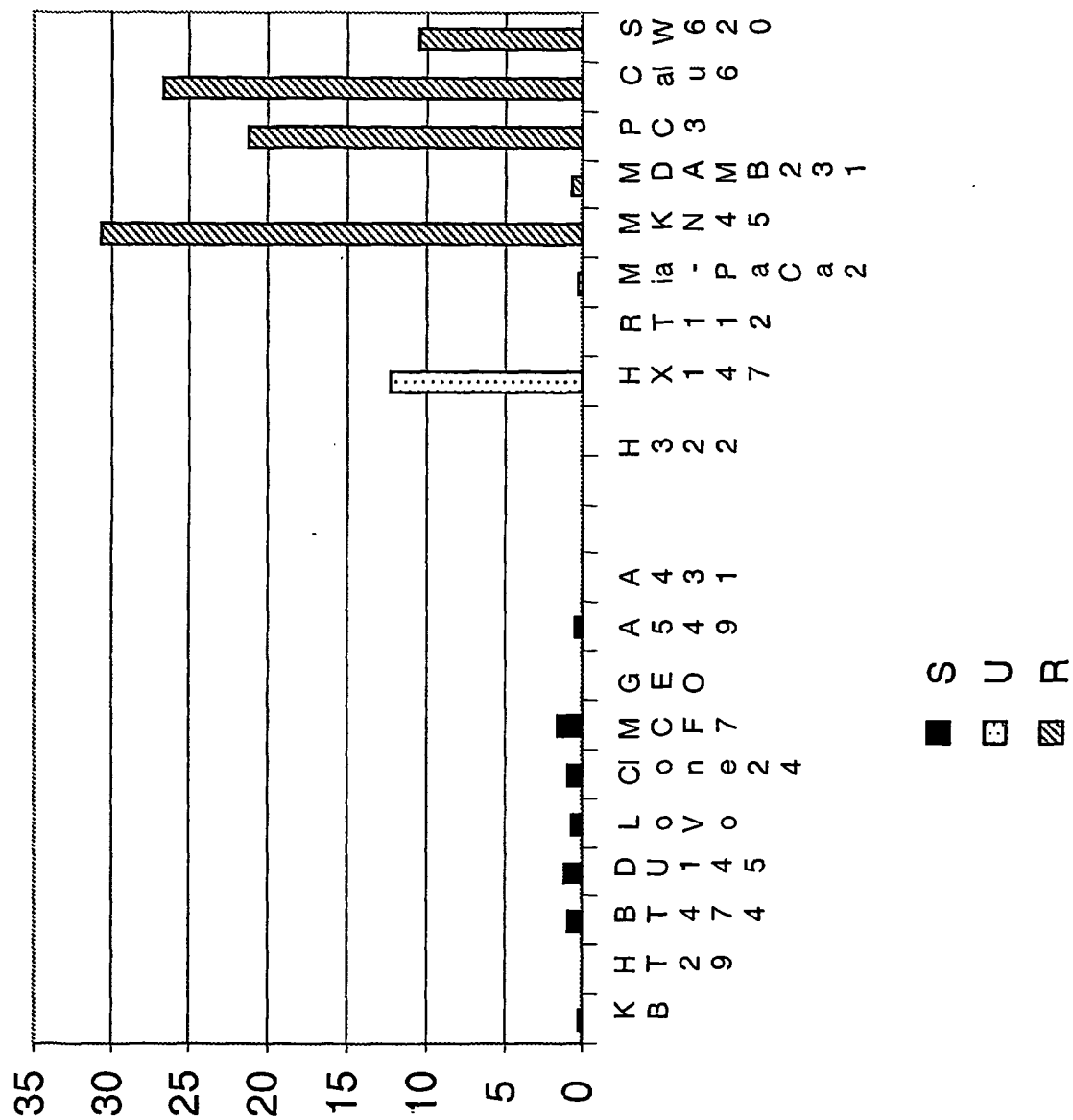


Figure 6



## 1

## SEQUENCE LISTING

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&lt;120&gt; METHOD

&lt;130&gt; CCH 101515

&lt;140&gt; US 60/590357 and US 60/619027

&lt;141&gt; 2004-07-23

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 gaggcaaatg gatctcgata tttcagatgg gcttttgatg cactgttgcc aaggaaggct 60  
 ttttctgatt ttttgacaaa tgaatttttg cacactttca ttggtgtctt tcggcaactt 120  
 acacacattg aaaat 135

<210> 7



## 3

<211> 402  
<212> DNA  
<213> Homo Sapiens

<400> 7  
caagttttgg tggcacgcag cctggggact ctgcctcgtg ccgctgagcc tggcgcagat 60  
cgatttgaat ataacctgcc gctttgcagg tgtattccac gtggagaaaa atggtcgcta 120  
cagcatctct cggacggagg ccgctgacct ctgcaaggct ttcaatagca ccttgcccac 180  
aatggcccag atggagaaaag ctctgagcat cggatttgag acctgcagtt tgcattgcag 240  
tcaacagtcg aagaagggtgt gggcagaaga aaaagctagt gatcaacagt ggcaatggag 300  
ctgtggagga cagaaagcca agtggactca acggagaggc cagcaagtct caggaaatgg 360  
tgcatttggt gaacaaggag tcgtcagaaa ctccagacca gt 402

<210> 8  
<211> 417  
<212> DNA  
<213> Homo Sapiens

<400> 8  
attgtaaatc ttttgtgtct cctgaagact tcccttaaaa ttagctctga gtgaaaaatc 60  
aaaagagaca aaagacatct tcgaatccat atttcaagcc tggtagaatt ggcttttcta 120  
gcagaacctt tccaaaagt ttatattgag attcataaca acaccaagaa ttgattttgt 180  
agccaacatt cattcaatac tgttatatca gaggagtagg agagaggaaa catttgactt 240  
atctggaaaa gcaaaatgta ctttaagaata agaataacat ggtccattca cctttatggt 300  
atagatatgt ctttgtgtaa atcatttggt ttgagtttcc aaagaatagc ccattgttca 360  
ttcttgtgct gtacaatgac cactgttatt gttactttga cttttcagag cacaccc 417

<210> 9  
<211> 546  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> misc\_feature  
<222> (104)..(104)  
<223> n is a, c, g, or t

<400> 9  
ttctatgcat ccacaccaa atcctgcaga atgtaagtaa gctctgcttt ataagatggg 60  
ttcaccttca tcgcagactg aaagtttcag tttttatttt tttncagaaa gcacgaaaaa 120  
ttatttataa tagtctggag aaaaaacaca ctgtaatat tcaagtgtat gcagtagaat 180  
gtactgtaac tgagcccttt ccacatgctc taggctccaa tgtctcctgt aggtccacct 240  
aactgtgtgt tttcaggagc aatgccatcc atgtttgtgc tgtagacttg ctgctgctga 300  
atcctttctg gggactttct catcgggcag ggagcagagg gcttctcggt catgcacct 360  
ttgcctgaac acccatgtag ctgctgtggt gtgtatatat tactcttaag aggagtgtgt 420  
gtgtctgtgt ttgttttaaa agtcacttat ttcttacagt gatttcaatt gcaccatgac 480  
ttcttcaact aaaccacaaa gtctgtctta aaactatgga aaacctaacc tgattagagc 540  
cttgac 546

<210> 10  
 <211> 546  
 <212> DNA  
 <213> Homo Sapiens

<400> 10  
 ggcaatctgt cacactctca gagtctggga cttgacttgc taccaacaac tgctgtgcaa 60  
 ttctgctgag caggaatatc atgagctgtt caataatgac ggacgcattg gttgagatga 120  
 agtttccagt aaggaagtga cagtgcattg tggatatatta tggctgtaaa ataggaagag 180  
 ctttagttcc caggctgaac ctgccactgc tggagccatt tcaacaaggc atcctcacia 240  
 caaagaagag atgtgatttg gtaccatttc acaccagcag gtgtctggac gaaaacatca 300  
 atgtgaataa gggccaagtg cagtcctgtc ttgattaaat tacttaataa tattattaaa 360  
 taataatagg tctgggcagt attgttttta acctgactca tccagctgtc cttcaaatag 420  
 ctccgtctcc ctctaccag aactgatttt taaaaagaag taatttttct ccctgggctg 480  
 ggaaaaccct aatgaactga aacacacttt tactttaaaa ttttctgtc tggcgttttt 540  
 gtaatc 546

<210> 11  
 <211> 496  
 <212> DNA  
 <213> Homo Sapiens

<400> 11  
 gaattcccta gaaatcctac tgggaagtat aggcagatct ctccctcata taacggatgt 60  
 ttcttggcgc ttggaatatc agataaagac caatcaactt cataggatgt acagacctgc 120  
 atatttgggtg accttaagtg tacagaacac tgattcccca tcctatccag agattagttt 180  
 tagttgcagc atggaacaat tacaggactt ggtggggaaa cttaaagatg cttcgaaaag 240  
 cctggaaaga gcaactcagt tgtaacttgg ggaagttaac gatccgcccg agtgcagagg 300  
 aaaaccagaa acgccttgcc ttcagctgaa ccaccgtttg tgcgagctgg atgtcctttt 360  
 cagtagaaaa gaattttcct tttgaattta taccattcat caattttgac actttaaaaa 420  
 cgtgtgaaag ggtaagagg gaaagatact gcccaagtat ttgaatcgtt tagtagtaac 480  
 tgtccattta tcctat 496

<210> 12  
 <211> 313  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> misc\_feature  
 <222> (190)..(190)  
 <223> n is a, c, g, or t

<400> 12  
 tataatactt cagtaaggcc tttaaaaaat ccacagtgat attattactc ctaacaaaaa 60  
 caataattac ttagtatcat ctaatatgtg gttcatattt aaatttggtg ttttgagatg 120  
 ggtcttacaa ttggtttatt caattgcatt ttttctaact cgtgtctcaa gtgttttaaa 180

## 5

aatctaactgn acttataatg acttatataa tgtattttctc attttacctt tttccaaaa 240  
gaggaaaataa tggcaaacca tataatatgtg tacatttcact gtcaaaaagc aaacccttgt 300  
tttgataact tgt 313

<210> 13  
<211> 395  
<212> DNA  
<213> Homo Sapiens

<400> 13  
cctcctccag ggtgatttta tgatcagtg tgttgcctc ggaagacatt tttccgtttg 60  
cttttgttcc aatgtcaatg tgaacgtcca catgaaacct acacactgtc atgcttcac 120  
attccctctc atctcaggta gaagggtgac acagttgtag ggttacagag acctatgtaa 180  
gaattcagaa gaccctgac tcacatttg tggcagtcct ttataattgg tgcatagcag 240  
atgggttcca catttagatc ctgggttcat aacttcctgt acttgaagtc taaaagcaga 300  
aaataaagga agcaagtttt cttccatgat tttaaattgt gatcgagttt taaattgata 360  
ggagggaaca tgcctaatt cttctgtcct gagaa 395

<210> 14  
<211> 569  
<212> DNA  
<213> Homo Sapiens

<400> 14  
aggagaggat ttgccactgc ttttctaagg acgagaagcc tgttgaagct attagggttt 60  
gttctgaagt ttacagatg gaacctgaca atgtgaatgc cctgaaagat cgagctgagg 120  
cctatttgat agaggaaatg tatgatgaag ctattcagga ttatgaaact gctcaggaac 180  
acaatgaaaa tgatcagcag attcgagaag gtctagagaa agcacaaaaga ttattgaaac 240  
agtcgcagaa acgagattat tataaaatct tgggagtaaa aagaaatgcc aaaaagcaag 300  
aaattattaa agcataccga aaattagcac tgcagtggca cccagataac ttccagaatg 360  
aagaagaaaa gaaaaaagct gagaaaaagt tcattgatat agcagctgct aaagaagtcc 420  
tctctgatcc agaaatgaga aagaagtttg acgacggaga agatcctttg gatgcagaga 480  
gccagcaagg agggggcggc aaccttttcc acagaagctg gaactcatgg caaggggttca 540  
atcccttcag ctcaggcgga ccatttaga 569

<210> 15  
<211> 481  
<212> DNA  
<213> Homo Sapiens

<400> 15  
tgagggccac gggcttgggt agtggaagg gtgtttggga aattgttaaa tcagttaccc 60  
gtagtagagc tatttcttgt acttctaagt tttctagaag tgggaaggatt gtagtcatcc 120  
tgaaaatggg ttacttcaa aatccctcag cctgttctt cactactgtc tatactgaga 180  
gtgtcatgtt tccacaaagg gctgacacct gagcctggat tttactcat cctgagaag 240  
ccctttccag taggggtggc aattcccaac ttccttgcca caagcttccc aggttttctc 300  
ccctggaaaa ctccagcttg agtcccagat acactcatgg gctgccctgg gcagccagca 360

## 6

```

ttcattgtaa gttccctctt tgaaaactgg tgtgtgggtg ttcagttctg tgtctgggtg 420
gtatggacag acagtaatct cctgtgatct gtgctagctg tgaggcagct ctggaacgtg 480
a 481

```

```

<210> 16
<211> 398
<212> DNA
<213> Homo Sapiens

```

```

<400> 16
ggctcccgagc aagggtagga cgggccgcat gcgggcagaa agttgggact gagcagctgg 60
gagcaggcga cagagctcct tccccatcat ttctccttgg ccaacgacga ggccagccag 120
aatggcaata aggactccga atacataata aaagcaaaca gaacactcca acttagagca 180
ataacggctg cgcagcagc cagggaagac cttggtttgg tttatgtgtc agtttcactt 240
ttccgataga aatttcttac ctcatTTTTT taagcagtaa ggcttgaagt gatgaaaccc 300
acagatccta gcaaagtgtc ccaaccagct ttactaaagg gggaggaagg gagggcaaag 360
ggatgagaag acaagtttcc cagaagtgcc tggttctg 398

```

```

<210> 17
<211> 499
<212> DNA
<213> Homo Sapiens

```

```

<400> 17
gatacgttgg ggcccatgca gaaggagctg gccgagcagc tgggcctgtc tactggcgag 60
aaggagaagc tgccgggaga gctagagccg gtgcaggcca cgcagaacaa gacagggaag 120
tatgtgccgc cgagcctgcg cgacggggcc agccgcgcgc gggagtccat gcagcccaac 180
cgcagagccg acgacaacgc caccatccgt gtcaccaact tgtcagagga cagcgtgag 240
accgacctgc aggagctctt ccggccttcc ggctccatct cccgcatcta cctggctaag 300
gacaagacca ctggccaatc caagggtttt gccttcatca gcttccaccg ccgagaggat 360
gctgcgcgtg ccattgccgg ggtgtccggc tttggctacg accacctcat cctcaacgtc 420
gagtgggcca agccgtccac caactaagcc agctgccact gtgtactcgg tccgggaccc 480
ttggcgacag aagacagcc 499

```

```

<210> 18
<211> 261
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (41)..(42)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (196)..(196)
<223> n is a, c, g, or t

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<400> 18
atgtgtcggg gagagagccc gcagggaagg gtaaagccca nnggggcagg gccctccag 60

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atgcctgagg aggggggcagg tccccctccc tctcctcctc tccccctccc atctaaagg 120  
 gtttggggag agacacaggc aggcgagggg gctggtcccc agtctgttgg ggtggtgctc 180  
 agggtaaagg gctatnggca acaggggacc agaccaggga tgagtgggga gggcacaagg 240  
 accatttgcc agaatccacc g 261

<210> 19  
 <211> 526  
 <212> DNA  
 <213> Homo Sapiens

<400> 19  
 ctgttgctcc aggatgcatt ctgataggag ggggcggcag ggctgggcct tgtgacaatc 60  
 tgccctttcac cacatggcct tgccctcgggt gccctgactg tcaggggagg ccaggagggc 120  
 agagcgggag ggagtctcag gaggaggcct gccctgaggg gctggggagg ggttacctca 180  
 tgaggaccag ggtggagcct gagaagagga ggaggtgggg gcttgagggt gcttggtagc 240  
 tgagggggac ggcaagttag aggggaggga ggggaagtcct gggaggatcc tgagctgctg 300  
 ttgcagtcta acccactaat cagttcttag attcagggga agggcaggca ccaacaactc 360  
 agaatggggg ctttcgggga gggcgccctag tccccccagc tctaagcagc caggaggggac 420  
 ctgcatctaa gcatctgggt tgccatggca atggcatgcc cccagctac tgtatgcccc 480  
 cgacccccgc agaggcagaa tgaacccata gggagctgat cgtaat 526

<210> 20  
 <211> 516  
 <212> DNA  
 <213> Homo Sapiens

<400> 20  
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 ccatcatccc cgagctggag cgtgagatcc gcatcatcaa cacggagcag tacatgcact 120  
 cgctgacgtg gcagcaggcg ctcacggggc tgctggagcg catgcagacc tatcaggacg 180  
 cggagtgcag gcagggtgctg gctgcctgga tgaaagagcg gcaggagctg aggtgcatca 240  
 ccaaggccct gttaatgcg cagttcggca gcatcttcg caccttccac aacccacct 300  
 acttctcaag gcgcctcgtg cgcttctctg acctctacat ggctccctc agctgcctgc 360  
 tcaactaccg cgtggacttc acctctacc cagcccgtag gccgctgcag cagagggcac 420  
 ccctctggat ggaccagctc tgcaccggct gcatgaagac ccccttcctt ggtgacatgg 480  
 cccacatccg ctgagggcac ctttattgtc tgggac 516

<210> 21  
 <211> 482  
 <212> DNA  
 <213> Homo Sapiens

<400> 21  
 tattcaaacg gagtccctccc attccaagaa actggaacc cctagtttat gttaaaaggc 60  
 cagtctaaat tctttcactt acatctttac agaaaactat atttctctc ttccataccc 120  
 agaaatctaa tcagaaaact gacttttctc atgttcaact ggacctaggg gaatatgaca 180

## 8

gaaaagcatc ccataggctt taatatactt tttaaaatat ataaaactga aaattaatag 240  
 ccatattaccc tgaaagagtt ctgctgggac tttgtcactt gcatagtaat agcatgtgcc 300  
 tcattgttca gaagattagc tttaggctct attttcaa atcgaaatgg agcataagct 360  
 gtaaaactgt agtcttctct gcagaaaata aaggccaaca ataagaaagc ttttgaagga 420  
 atcacggaaa acaaatttat aaaagaaata actatatgcg cagtaattct taacacattg 480  
 ac 482

<210> 22  
 <211> 459  
 <212> DNA  
 <213> Homo Sapiens

<400> 22  
 gcaagtcgcg tgatttctac cacacctgct actgcctgag cggcctgtcc atagcccagc 60  
 acttcggcag cggagccatg ttgcatgatg tggctcctggg tgtgcccga aacgctctgc 120  
 agcccaactca cccagtgtac aacattggac cagacaaggt gatccaggcc actacatact 180  
 ttctacagaa gccagtccea ggttttgagg agcttaagga tgagacatcg gcagagcctg 240  
 caaccgacta gaggacctgg gtcccggcag ctctttgctc acccatctcc ccagtcagac 300  
 aaggtttata cgtttcaata catactgcat tctgtgctac acaagcctta gcctcagtgg 360  
 agctgtgggt ctcttggtac tttctgtca aaaaaacca atggctctgg gtttgagaa 420  
 cacagtggct ggttttaaaa ttctttccac acctgtcaa 459

<210> 23  
 <211> 549  
 <212> DNA  
 <213> Homo Sapiens

<400> 23  
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 cctcacaagc agtgacacct cgggtccttt ccgttgctat ggtgaaaatt cctggatgga 120  
 atggatcaca tgagggttcc ttgttgcttt tggagggtgt gggggatatt ttgttttgg 180  
 ttttctgcag gttccatgaa aacagccctt ttccaagccc attgtttctg tcatggtttc 240  
 catctgtcct gagcaagtca ttctttgtt atttagcatt tcgaacatct cggccattca 300  
 aagcccccat gttctctgca ctgtttggcc agcataacct ctagcatcga ttcaaagcag 360  
 agttttaacc tgacggcatg gaatgtataa atgagggtgg gtccttctgc agatactcta 420  
 atcactacat tgctttttct ataaaactac ccataagcct ttaaccttta aagaaaaatg 480  
 aaaaagggtta gtgtttgggg gccgggggag gactgaccgc ttcataagcc agtacgtctg 540  
 agctgagta 549

<210> 24  
 <211> 372  
 <212> DNA  
 <213> Homo Sapiens

<400> 24  
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 cacctactct tatttactgc ccattgattg acttttcttc atattttgca aagagaaatt 120

tcacagcaaa aattcatgtt ttgtcagctt tctcatgttg agatctgtta tgtcactgat 180  
 gaatttaccc tcaagtttcc ttctctgtga ccactctgct tccttggaaca atatcagtaa 240  
 tagctttgta agtgatgtgg acgtaattgc ctacagtaat gaaaaattaa tgtacttta 300  
 tttttcattt tcttttagga tatttagacc acccttggtc caccgcaaac agagtgtgtc 360  
 agtgtttgtg tg 372

<210> 25  
 <211> 475  
 <212> DNA  
 <213> Homo Sapiens

<400> 25  
 cagggatcgg aggacgaccc gagtcccaag agtgggggtt tgcttttttaa aaggagagag 60  
 gaggggtgat ggcaggggag tggaggggtg ccgggcaggc cctgccggcg cagggagccc 120  
 tctgcccttc acactctcct ccaaaagagc ctccatctgt aaggagcag gtctccgga 180  
 ggggtttctt tccatgtgtt ttctctctgt tgttaaaaga acttttttaa aaaaacagac 240  
 ctctgttttag atttatagca ttgactttta cacacattca cacaagaaaa aaatccttc 300  
 aaaattctta aatcttctgt tctctctttt tccaagggaa gagggcaaaa agtggcctgg 360  
 gctctgttgg tgtgcgtgtt ccgtggcgga gagaagaaaa tgggaaagac atctcactgg 420  
 tgcttttctc ttttgtttta gtgcccccg ccccatccc tataatatct gtaac 475

<210> 26  
 <211> 516  
 <212> DNA  
 <213> Homo Sapiens

<400> 26  
 gaagcaattg ctcatgttgg ccaaacatgg tgcaaccagt gatttccatc tctggtaaag 60  
 ttacactttt atttctgtga tgtgtacaa tcaaaacaca ctactacctc ttaagtccca 120  
 gtatacctca tttttcatac tgaaaaaaaa agcttgtggc caatggaaca gtaagaacat 180  
 cataaaattt ttatatatat agtttatttt tgtgggagat aaattttata ggactgttct 240  
 ttgctgttgt tggctgcagc taaataagac tggacattta acttttctac catttctgca 300  
 agttaggtat gtttgccagg agaaaagtat caagacgttt aactgcagtt gactttctcc 360  
 ctgttccttt gagtgtcttc taactttatt ctttgttctt tatgtagaat tgctgtctat 420  
 gattgtactt tgaatcgctt gacttgttga aaatatttct ctagtgtatt atcactgtct 480  
 gttctgcaca ataaacataa cagcctctgt gatccc 516

<210> 27  
 <211> 566  
 <212> DNA  
 <213> Homo Sapiens

<400> 27  
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 taagcggta tcgttgcgtc atggggcagg cgtggggagc ttctgtctgc cttggctggg 120  
 tgtgggcctg gaggaaggtc ctggggcgtg cactgcctg ggcagtgggg aggagagtgg 180

## 10

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cctgagttac ttcacccccg cgtgctgctg gttaatgtcc cgcgtctctg caccttcggg 240
tgaggagcggg gactgatcta ctttcacatt ctcaagtttt tctcatctgc attagaggtc 300
cccagtaggt tcccagggtc cagcgtgccc ctccctcaga cacacggaca caatcagccg 360
agaagttcct ggtctgaatc acgagaatgt ggaggggtgg ggggtgtcag tggaaaggca 420
taaggctgag ctgagaccag ttgctgggtga aactgggcca atctggggag gggaacatcc 480
ttgccaggga gtttctgagg gtctgctttg tttacctttc gtgcggtgga ttctttttaa 540
ctccgtctac ctggcgtttt gttaga 566

```

```

<210> 28
<211> 327
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> misc_feature
<222> (199)..(199)
<223> n is a, c, g, or t

```

```

<400> 28
ccacctgtga ccccggtgtg gaggagcatt tccgcaggag cctgggcaag aattacaagg 60
agccccagcc ggcaaccaac tccgtgtcca tcacgggctc cgtggacgac cactttgcc 120
aagctctggg tgacacgtgg ctccagatca aagcggccaa ggacggagca tccagcagcc 180
ctgagtcgcc ctctcgcan ggcagccccg ccagcccctc tgcccacatg gtcagccaca 240
gtcactcccc ctctgtgtgc tctgaaggg agcgccctct ccaacaacac gtggatctgc 300
atggtttgcc tgagctttga acagtca 327

```

```

<210> 29
<211> 347
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> misc_feature
<222> (156)..(156)
<223> n is a, c, g, or t

```

```

<400> 29
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tttttacttc cttctgggtg gaggttttcc gactcccaat catgaaggca agttaatctt 120
tccagttagt gacttttgcc ccatagttgg ggtaancact tctagattg agaaaaagca 180
gctacagtca atcctgctct gtttgctca tttggtgatc agtcagtcac acataagtcc 240
cttgtattct aaatttcatg cacttctccc agatgctata gggttttctc tcactgttgc 300
caatggatgt catccagaca gtgggctcat atcttacggg tttgtgc 347

```

```

<210> 30
<211> 210
<212> DNA
<213> Homo Sapiens

```

```

<400> 30
agttgatcag agccttccag agtgtggtat gcttttcaact gtgtgatgat ccttagtggc 60

```



## 11

acatgaatga acgtccagat gtttgtgcag tagccacccc ttatctgcag gatacgttcc 120  
aagacccccca gtgaatgcct gaaactgcag atagtactga atcctatata tactgtgttt 180  
tttatgatac atacatgcct atgatgaagt 210

<210> 31  
<211> 511  
<212> DNA  
<213> Homo Sapiens

<400> 31  
aagagaatgt tcctactcac acttcagctg ggtcacatcc atccctccat tcatccttcc 60  
atccatcttt ccatccatta cctccatcca tccttccaac atatatttat tgagtaccta 120  
ctgtgtgccca ggggctggtg ggacagtggg gacatagtct ctgccctcat agagttgatt 180  
gtctagttag gaagacaagc attttttaaaa aataaattta aacttacaaa ctttgtttgt 240  
cacaagtggg gtttattgca ataaccgctt ggtttgcaac ctctttgctc aacagaacat 300  
atgttgcaag accctcccat gggggcactt gagttttggc aaggctgaca gagctctggg 360  
ttgtgcacat ttctttgcat tccagctgtc actctgtgcc tttctacaac tgattgcaac 420  
agactgttga gttatgataa caccagtggg aattgtctga ggaaccagag gcacttccac 480  
cttggtctggg aagactatgg tctgccttg c 511

<210> 32  
<211> 505  
<212> DNA  
<213> Homo Sapiens

<400> 32  
aaggcattcc acaggatcat catttaaaaa aaaagaattc tggctcctgtt ttctaaaaaa 60  
aaaaaactgt tgtagaaatt cttaatttgg atctatttat tagtcagagt ttcagctttc 120  
ttcagctgcc agtgtgttac tcatctttat cctaaaaatc tggaatcaga gatttttgtt 180  
tgttcacata tgattctctt agacactttt atatttgaaa aaattaaaaat ctttctttgg 240  
ggaaaaatcc ttggttatcc tgccataaca gattatgtat taactttagat attcagtggg 300  
tcaataacctg tttagtgtgt tgctaataatt tccagaagga tttcttgtat tggtgaaaga 360  
cggttgggga tgggggggatt tttttgttct tgttgtacct ttgttttgaa actagaaatc 420  
tgtcctgtgg catgcaaaag aaagcaaatt atttttaaaa gaaaaaaacc aaagtacttt 480  
tggtgtcatt attccatctt ctcca 505

<210> 33  
<211> 307  
<212> DNA  
<213> Homo Sapiens

<400> 33  
ccagccactg cagatagaga catatggacc acatgttcct gagcttgaga tgctaggaag 60  
acttgggtat ttaaaccatg tcagagctgc ctaccacag gaccttgctg gaggtatac 120  
ttcttctctt gcttgcaca gagcactaca ggatgcattc agtgggcttt tctggcagcc 180  
cagttaacca ttataagat ttggaccttg gagctgaacc agggagctag caaaagtaaa 240

## 12

gcagacttat aaaattatag ctatgtgcag ctgcacaaca cagtccttcc actagcagct 300  
gtgttaa 307

<210> 34  
<211> 519  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> misc\_feature  
<222> (130)..(130)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (144)..(144)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (167)..(167)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (169)..(169)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (268)..(268)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (349)..(349)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (358)..(358)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (478)..(478)  
<223> n is a, c, g, or t

<400> 34  
caccgcgcag agctcagggg gtggtgcgcc cggcccttct gcggcgcaaca gccagccca 60  
ggaacgcggg cggtgcggac tcagcgggcc ggggtgcaggc gcggagctgg gcctctgcgc 120  
ccggcccgan ctccgtctat aaanagagca gccagttgca gggctcnant ctgctttcca 180  
actgcctgac tgcttggtcg tctcactggt gtgagctoca gcatccctt tgctcgaaat 240  
ggaccccaac tgctcttgcg ccaactggntg gctcctgcac gtgcgccggc tcctgcaagt 300  
gcaaagagtg caaatgcacc tcctgcaaga agagctgctg ttctgtctnc ccctgggnct 360  
gtgccaagtg tgcccagggc tgcgtctgca aaggggcac gcgagaagtgc agctgctgtg 420  
cctgatgtgg gaacagctct tctcccagat gtaaataagaa caacctgcac aacctggnat 480  
ttttttaaaa atacaacact gagccatttg ctgcatttc 519

<210> 35

## 13

<211> 460  
 <212> DNA  
 <213> Homo Sapiens

<400> 35  
 gcagcactct taacttacga tctcttgaca tacggtttct ggctgagagg cctggcccgcc 60  
 taagggtgaaa aggggtgtgg caaaggagcc tactccaaga atggaggctg taggaatata 120  
 acctcccacc ctgcaaaggg aatctcttgc ctgctccatc tcataggcta agtcagctga 180  
 atcccgatag tactagggtcc ccttccctcc gcatcccgtc agctggaaaa ggctgtgtgc 240  
 ccagaggctt ctccaaaggg aggggtgacat gctggctttt gtgccaagc tcaccagccc 300  
 tggccacact cactgcagta gtgcaccatc tcaactgcagt agcacgccct cctgggcccgt 360  
 ctggcctgtg gctaattggag gtgacggcac tcccatgtgc tgactcccc catccctgcc 420  
 acgctgtggc cctgcctggc tagtccttgc ctgaataaag 460

<210> 36  
 <211> 540  
 <212> DNA  
 <213> Homo Sapiens

<400> 36  
 gctacagatt cacactttct ggcctaaacc ctaatgggat gaggttttcc accccaggcc 60  
 atgctgggtg tgatttttta gcccctaaat aaaacactgg actatttccct gtttacttca 120  
 ttgattgcaa ctacaaagggt ggactcaaag caaagcacia tcatgccagc caacattcca 180  
 gaattctgct gagaactcca agtctgtgag gggagagggt ttacaagcca gacaggcctg 240  
 ggggactgca gtccccaagg agaccctgcc acatgctggc cctttgagtg agaattgctgc 300  
 atctttctac atatcttcat gagaatactg agaattggat tttccttttc aaaatgcact 360  
 ttgctttttt tgtatgtttt gttatgttga gatgtttcta aagaaaagat tttatgtaat 420  
 tataagatga agcgtagtga attgtacagc tgttgaata atgacctatt tctatataaa 480  
 ataaaattgt atggcttatg tgtaaattat tttgtatctg agataccagt tccttttccc 540

<210> 37  
 <211> 367  
 <212> DNA  
 <213> Homo Sapiens

<400> 37  
 aaaggggatg gacgtctcat tctcagggat cctgtctttc attgaggatg tagcccatcg 60  
 gatgctggcc acaggcgagt gtactcctga ggatctgtgt ttctccctgc aggaaactgt 120  
 gtttgcaatg ctggtagaga tcacagagcg agccatggca cattgtggct ccaggaggcc 180  
 cctcattgtg ggaggagtgg ggtgtaattgt gaggctacag gagatgatgg caacaatgtg 240  
 ccaggaacgt ggagcccggc tttttgctac agatgagaga ttctgtattg acaatggagc 300  
 gatgatagcc caggctggct gggagatggt tcgggctgga cacaggaccc cactcagtga 360  
 ttctggg 367

<210> 38  
 <211> 532  
 <212> DNA  
 <213> Homo Sapiens

## 14

<400> 38  
 cagaaagtct cagcccagga tggggcttct tcaacagggc cctgccctc ctgaagcctc 60  
 agtccttcac cttgccaggt gccgtttctc ttccgtgaag gccactgcc aggtccccag 120  
 tgcgccccct agtggccata gcctgggtaa agttccccag tgctccttg tgcataagacc 180  
 ttctttctcc acccccttct gccctgggt ccccgccat ccagcggggc tgccagagaa 240  
 cccagacct gcccttacag tagtgtagcg cccctccct ctttcggctg gtgtagaata 300  
 gccagtagtg tagtgcggtg tgcttttacg tgatggcggg tgggcagcgg gcggcgggct 360  
 ccgcgcagcc gtctgtcctt gatctgccg cggcgggccg tgtgtgttt tgtgtgtgt 420  
 ccacgcgcta aggcgacccc ctccccgta ctgacttctc ctataagcg ttctcttcgc 480  
 atagtcaagt agtcccacc ccacctctt cctgtgtctc acgcaagttt ta 532

<210> 39  
 <211> 551  
 <212> DNA  
 <213> Homo Sapiens

<400> 39  
 ggatggggct tcttcaacag ggccccctgc ctctgaagc ctcaagtcct cacttgcca 60  
 ggtgccgttt ctcttcctg aaggccactg ccaggtccc cagtgcgccc cctagtggcc 120  
 atagcctggt taaagttccc cagtgcctcc ttgtgcatag acctcttct cccaccccct 180  
 tctgccctg ggtccccggc catccagcgg ggctgccaga gaaccccaga cctgccctta 240  
 cagtagtgta gcgccccctc cctctttcgg ctggtgtaga atagccagta gtgtagtgcg 300  
 gtgtgctttt acgtgatggc ggggtgggag cggcgggcgg gctccgcgca gccgtctgtc 360  
 cttgatctgc ccgcgccggc ccgtgttggt tttgtgtgt tgtccacgcg ctaaggcgac 420  
 cccctcccc gtactgactt ctccataag cgcttctct cgcatagtca cgtagctccc 480  
 accccacct ctctgtgt ctcaagcaag ttttatact taatatatt atggcttttt 540  
 ttcttcgaca a 551

<210> 40  
 <211> 538  
 <212> DNA  
 <213> Homo Sapiens

<400> 40  
 gccagctttg ggctgagcta acaggaccaa tggattaaac tggcatttca gtccaaggaa 60  
 gctcgaagca ggtttaggac caggtccct tgagaggtca gaggggcctc tgtgggtgct 120  
 gggtagtcca gaggtgccac tgggtggaagg gtcagcggag cccagtgcc tcttgtgca 180  
 tagac-ttct tctccacccc cttctgccc ctgggtccc gccatccag cggggtgcc 240  
 agagaacccc agacctgcc ttacagtagt gtacgcccc ctccctctt cggctggtgt 300  
 agaatagcca gtagttagt gcggtgtgt tttacgtgat ggcgggtggg cagcggcgg 360  
 cgggctccgc gcagccgtct gtccttgatc tgcccgggc ggcccggtgt gtgttttgtg 420  
 ctgtgtccac gcgctaaggc gacccctcc cccgtactga cttctctat aagcgttct 480  
 cttcgcatag tcacgtagct cccacccac cctcttctg tgtctcagc aagtttta 538

## 15

<210> 41  
 <211> 403  
 <212> DNA  
 <213> Homo Sapiens

<400> 41  
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 ccatccggaa cagggccttc tacatggagg agggcgtgcc ctattgagag cgagactatg 120  
 agaagatgtt tggcacgaaa tgccatggct gtgacttcaa gatcgacgct ggggaccgct 180  
 tcctggaggc cctgggcttc agctggcatg acacctgctt cgtctgtgag atatgtcaga 240  
 tcaacctgga aggaaagacc ttctactcca agaaggacag gcctctctgc aagagccatg 300  
 ccttctctca tgtgtgagcc ccttctgccc acagctgccg cgggtggcccc tagcctgagg 360  
 ggcttgaggt cgtggccctg catttctggg tagggtggc aat 403

<210> 42  
 <211> 437  
 <212> DNA  
 <213> Homo Sapiens

<400> 42  
 tggatccaaa ctttattat gccattatat gatgccagat gaagaaactc cattagcagt 60  
 gcaggcctgt ggactttctc ctcgagacat taccactatt aaacttctca atgaaactag 120  
 agacatgttg gaaagcccag atttttagtac agttttgaat acctgtttaa accgagggtt 180  
 tagtagactt ctagacaata tggtgagtt ctttcgacct actgaacagg acctgcaaca 240  
 tggtaactct atgaatagtc tttccagtgt cagcctgcct ttagctaaga taattccaat 300  
 agtaaacgga cagatccatt cagtttgcag tgaacacct agtcattttg ttcaggatct 360  
 gttgacaatg gagcaagtga aagactttgc tgctaagtgt tatgaagctt ttagtaccac 420  
 tcagcaactg gagaaat 437

<210> 43  
 <211> 520  
 <212> DNA  
 <213> Homo Sapiens

<400> 43  
 agccatttga agacgcctcg tttgcgctgc ggacggggga gatgagcggg cccgtgttca 60  
 cggattccgg catccacatc atcctccgca ctgagtgagg gtggggagcc caggcctggc 120  
 ctcggggcag ggcagggcgg ctaggccggc cagctcccc ttgcccgcga gccagtggcc 180  
 gaacccccca ctccctgcca ccgtcacaca gtattttattg ttcccacaat ggctgggagg 240  
 gggcccttcc agattggggg ccctggggtc cccactccct gtccatcccc agttggggct 300  
 gcgaccgcca gattctccct taaggaattg acttcagcag ggggtgggagg ctcccagacc 360  
 cagggcagtg tgggtgggagg ggtgttccaa agagaaggcc tggtcagcag agccgccccg 420  
 tgtcccccca ggtgctggag gcagactcga gggccgaatt gtttctagtt aggccacgct 480  
 cctctgttca gtcgcaaagg tgaacactca tgcggcagcc 520

<210> 44  
 <211> 530

## 16

<212> DNA  
 <213> Homo Sapiens

<220>  
 <221> misc\_feature  
 <222> (68)..(68)  
 <223> n is a, c, g, or t

<400> 44  
 gattaaacga ctgtgtcttt gtcacctctg cttaacttta ggagtatcca ttctgtgat 60  
 tgtagacntt tgttgatatt cttcctggaa gaatatcatt cttttcttga agggttgggt 120  
 tactagaata ttcaaatca atcatgaagg cagtactat tttagtcta aaggttttct 180  
 aaaaattaac ctacatccc ttctgttagg gtctttcaga atatctttta taaacagaag 240  
 catttgaagt cattgctttt gctacatgat ttgtgtgtgt gaaggacata ccacgtttta 300  
 atcattaatt gaaaaacatc atataagccc caactttgtt tggaggaaga gacggagggt 360  
 gaggtttttc cttctgtata agcacctact gacaaaatgt agaggccatt caaccgtcaa 420  
 acaccatttg gttatatcgc agaggagacg gatgtgtaaa ttactgcatt gctttttttt 480  
 tcagtttgta taacctctaa tctccgttg catgatacgc tttgttagaa 530

<210> 45  
 <211> 485  
 <212> DNA  
 <213> Homo Sapiens

<400> 45  
 tgaatgtacg cttgtccatg ctgacctcag tgagtataac atgctgtggc atgctggaaa 60  
 ggtctggttg atcgatgtca gtcagtcagt agaacctacc caccctcacg gcctggaggt 120  
 cttgttccgg gactgcagga atgtctcgca gtttttccag aaaggaggag tcaaggaagc 180  
 ccttagtgaa cgagaactct tcaatgctgt ttcaggctta aacatcacag cagataatga 240  
 agctgatttt ttagctgaga tagaagcttt ggagaaaatg aatgaagatc acgttcagaa 300  
 gaatggaagg aaagctgctt cttttttgaa agatgatgga gaccaccac tactatatga 360  
 tgaatagcac taataccac tgcttcagt ttaacacagc agtgattgtc agtgccaat 420  
 agcaaatgaa gttatgggtg acttgaaata ccaaacctg aggagtgggc aatggtgctt 480  
 ctgtg 485

<210> 46  
 <211> 351  
 <212> DNA  
 <213> Homo Sapiens

<400> 46  
 ttgcgccat tggccgtgtt ggtcttgaac tcttgccctc aagcaatccg cctacctcag 60  
 cctcccaaag tgctaggatt acaggcataa gccactgagc ccagccctag ttcagtatct 120  
 tttatgtaaa ttataaacat ctgcaacatt atgtatcata tgcagatact tattgcattt 180  
 cttttattag tgggtgaaagt gttctatgca tttattggct cttgaatttc ctcatctatg 240  
 aattgtcatt cacacaccta cttttctgct tcgtttttac atatgtcttt gcctattaaa 300  
 gatattatcc ctctgtttta tttttctct cattcttgta ttgcctttta a 351

## 17

<210> 47  
 <211> 521  
 <212> DNA  
 <213> Homo Sapiens

<400> 47  
 ccggaggcaa agagaccggg ccgaggaggc caaggaaagg gagaacaccg aaaacaataa 60  
 ctctctctcc aacaagcaga accaactctc tcctctggaa gggggcaagc cgctcatgtc 120  
 cagctcagaa gaggaattct cacctcccca aagtccagac cagaactcgg tccttctgct 180  
 gcagggcaat atggggccag ccaggagctc aaactattct ctcccgggct taacagcctc 240  
 gcagcccagt cacggcctgc agaccaccca gcatcagctc caagactctc tgctcgggcc 300  
 cctcacctcc agtctgggtg acttggggtc ctaagtgggg agggactggg gcctcgaagg 360  
 gattctctga gcagcaacca ctgcagcgac tagggacact tgtaaataga aatcaggaac 420  
 atttttgcag cttgtttctg gagttgtttg cgcataaagg aatggtggac tttcacaaat 480  
 atctttttta aaatcaaaac caacagcgat ctcaagctta a 521

<210> 48  
 <211> 498  
 <212> DNA  
 <213> Homo Sapiens

<400> 48  
 ggctgagcac cagtgaagtc tttgcctcta ctctgacct agacaacctg gggaggggacc 60  
 ctgtgcccgc aaaccagaca cataggacaa agtttatcta taacctggaa gaccatgagt 120  
 ggtgtgaaaa catggagtc gttttatagt gactaaagga gggctgaact ctgtattagt 180  
 aatccaaggg tcattttttt cttaaaaaaa gaaaaaagg ttccaaaaaa aacaaaaact 240  
 cagtacacac acacaggcac agatgcacac acacgcagac agacacaccg actttgtcct 300  
 ttttctcagc atcagagcca gacaggattc agaataagga gagaatgaca tcgtgcggca 360  
 gggctctgga ggccactcgc gcggctgggc cacagagtct actttgaagg cacctcatgg 420  
 ttttcaggat gctgacagct gcaagcaaca ggactgcca aattcagga acagtgggtg 480  
 ccagcttgga ggatggac 498

<210> 49  
 <211> 331  
 <212> DNA  
 <213> Homo Sapiens

<400> 49  
 gagacgtggg aagtgcggtg cagttttcaa ctgacctctg gacgcagaac ttcagccatg 60  
 aaggtaacag gcatctttct tctcagtgcc ttggccctgt tgagtctatc tggtaacact 120  
 ggagctgact ccctgggaag agaggccaaa tgttacaatg aacttaatgg atgcaccaag 180  
 atatatgacc ctgtctgtgg gactgatgga aatacttate ccaatgaatg cgtgttatgt 240  
 tttgaaggtc ggaaacgcca gacttctatc ctcatcaca aatctggggc ttgctgagaa 300  
 ccaaggtttt gaaatcccat caggtcaccg c 331

<210> 50  
 <211> 548

## 18

<212> DNA  
 <213> Homo Sapiens

<400> 50  
 agccatccca tgtagagct tctcaagagg aagacagccc agactctttc agttctctgg 60  
 attctgagat gtgcaaagac taccgagtat tgcccaggat aggetatctt tgtccaaagg 120  
 atttaaagcc tgtctgtggt gacgatggcc aaacctacaa caatccttgc atgctctgtc 180  
 atgaaaacct gatacgccaa acaatacac acatccgcag tacaggaag tgtgaggaga 240  
 gcagcacccc aggaaccacc gcagccagca tgccccgctc tgacgaatga caggaagatt 300  
 gttgaaagcc atgagggaaa aaataaaccc cagttctgaa tcacctacct tcaccatctg 360  
 tatatacaaa gaattcttcg gagcttgtct tatttgctat agaaaacaat acagagcttt 420  
 tgggaatgga atcactgatt ttcagtcttt tccatttctt tctcctaga atctgtgatc 480  
 tgaggggtata aagacatttc caccaagttt gagccctcaa aatgtcctga ttacaatgct 540  
 gtctgtcc 548

<210> 51  
 <211> 526  
 <212> DNA  
 <213> Homo Sapiens

<400> 51  
 gtccacattc ctgcaagcat tgattgagac atttgacaaa tctaaaatgt aagcaaagta 60  
 gtcattaaaa atacaccctc tacttgggct ttatactgca taaaaattta ctcatgagcc 120  
 ttcctttgag gaaggatgtg gatctccaaa taaagattta gtgtttattt tgagctctgc 180  
 atcttaacaa gatgatctga acacctctcc tttgtatcaa taaatagccc tggtattctg 240  
 aagtgagagg accaagtata gtaaaatgct gacatctaaa actaaataaa tagaaaacac 300  
 caggccagaa ctatagtcac actcacacaa agggagaaat ttaaaactga accaagcaaa 360  
 aggcttcacg gaaatagcat ggaaaaacaa tgcttcaggt ggccacttcc taaggaggaa 420  
 caaccccgctc tgatctcaga attggcacca cgtgagcttg ctaagtgata atatctgttt 480  
 ctactacgga tttaggcaac aggacctgta cattgtcaca ttgcat 526

<210> 52  
 <211> 476  
 <212> DNA  
 <213> Homo Sapiens

<400> 52  
 tgggggactt atttgttggg gatcttaaat aagattcctt ttgatctacc ggaatatata 60  
 tgtacagagt acattggatc atgttggaaa gaaggcaagt gaaaaggta gagatgaagt 120  
 agcgaagtta tggaatatcg tggaaaggat actagtgtg aaatggaaag agacaagttt 180  
 tagtacccca aaagcaaac aagcaggaga tgcaagagat gccccaaaag gacaaagcaa 240  
 caattttctg ttgccacctt tataccggaa gactctgttg tagaagaaaa gaaggctttg 300  
 gtgcacctta tgtgggagga ggaggggagc ggcattgctga tgctgagcgt acaggcagac 360  
 aagagcgtag cctgctgttg cctccatcac tatgaaatga cttattttac ctgaaggacc 420  
 catggtttat gtctctctaa ttcctttcac tctccctaag ccctctgaga gagatg 476



## 19

<210> 53  
 <211> 501  
 <212> DNA  
 <213> Homo Sapiens

<400> 53  
 gcctgtcggc tcagatcgag gaatgcatct tccgggacgt tggaaacaca gacatgaagt 60  
 ataagaaccg tgtacggagt cgtatctcca acctgaagga tgccaagaac cctgacctgc 120  
 ggcggaatgt gctgtgtggg gccataacac cccagcagat cgctgtgatg acctcagagg 180  
 agatggccag tgatgagctg aaggagatcc gtaaggccat gaccaaggag gccatccgag 240  
 agcaccagat ggcccgcact ggccgcacgc agacagacct gttcacctgc ggcaagtgc 300  
 ggaaaaagaa ctgcacctac acacaggtgc agaccgcag ctctgatgag cccatgacca 360  
 cctttgttgt ctgcaacgag tgtggaaacc gctggaagtt ctgctgacct ctcgtgtaga 420  
 tgtgtctgcag ccttggggccc tccccggccc acgtcctccg ttgacacagc ttctctggag 480  
 accctagaag gggcatgtc c 501

<210> 54  
 <211> 453  
 <212> DNA  
 <213> Homo Sapiens

<400> 54  
 tggatgacat ctacaaggct gcggtagagc agctgacaga agagcagaaa aatgagttca 60  
 aggcagcctt cgacatcttc gtgctgggcg ctgaggatgg ctgcatcagc accaaggagc 120  
 tgggcaagyt gatgaggatg ctgggcccaga accccacccc tgaggagctg caggagatga 180  
 tcgatgaggt ggacgaggac ggcagcggca cgggtggactt tgatgagttc ctggtcatga 240  
 tggttcggtg catgaaggac gacagcaaag ggaaatctga ggagctgtct gacctcttcc 300  
 gcatgtttga caaaaatgct gatggctaca tcgacctgga tgagctgaag ataatgctgc 360  
 aggctacagg cgagaccatc acggaggacg acatcgagga gctcatgaag gacggagaca 420  
 agaacaacga cggccgcatac gactatgatg agt 453

<210> 55  
 <211> 498  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> misc\_feature  
 <222> (142)..(142)  
 <223> n is a, c, g, or t

<400> 55  
 acccttgccc atcaggcgag gggctgggccc tgtgcagctg ggcccttggc cagagtccac 60  
 tcccttccctg gctgtgtcac cccgagcagc tcatccacca tggaggtcat tggcctgagg 120  
 caagttcccc ggagagtcgg gntccctgt gggccctca ggcctatgtc tgtgaggaag 180  
 gggccctgcc actctcccca agagggcctc catgtttcga ggtgcctcaa catggagcct 240  
 tgcttgccct gggctagggg cactgtctga actcctgact gtcaggataa actccgtggg 300  
 ggtacaggag cccagacaaa gccaggcct gtcaagagac gcagagggcc cctgccaggg 360

## 20

ttggccccag ggaccctggg acgaggctgc agaagctctc cctccctact cctggggagc 420  
 cacgtgctgg ccatgtggcc agggacggca tgagcaggag gcggggacgt gggggccttc 480  
 tggtttggtg tcaacagc 498

<210> 56  
 <211> 544  
 <212> DNA  
 <213> Homo Sapiens

<400> 56  
 gaagccacaa agatgccaca tgtagtata tcagtggagag gtgactccac agtgctctct 60  
 ggagaagcaa tatgagtgc tgaagagtgg ggctttttgc ttttgcctgg atataggggt 120  
 gctctcttac tgtaattggg tgtggaaaaa ctctggcttt atggtattcc attaggttct 180  
 tttcatttaa agtagtctta aaatcaaagt atccaatatt ttaaagccac aaagtagatt 240  
 acataattag cagagatttt agtcagtaaa atgttagaaa tcaaaactata agaaaattca 300  
 agtcccttat tttgtgtctt gggatatatgt cattatttta aattccacac tcccttattt 360  
 aatcactttg gtaagtgcct ttgatgtttt gaaatgtata gtgggagatg agcaaagtga 420  
 aatgtcatgt gccctgttcc ctagcttctc aattcctcat aaccattttt accagtgttg 480  
 caaagtttag acctttgtgt taatatcaga agtgtatttg tagccctcc atagtgaaca 540  
 atga 544

<210> 57  
 <211> 535  
 <212> DNA  
 <213> Homo Sapiens

<400> 57  
 gccgaagaag cctgtctgtg ggggtgtgtc cagcgctctg gcacctggcg tccgagccgt 60  
 ggagctcgag cggcagatcg agagcacaga gacttcttgc catggctgcc gtaagaattt 120  
 ctctctgtcc aagatccggt cccacgtggc tacttgttcc aaataccaga attacatcat 180  
 ggaagggtgtg aaggccacca ttaaggatgc atctcttcag ccaaggaatg tcccaaaccg 240  
 ttacaccttt ccttgcctt actgtcctga gaagaacttt gatcaggaag gacttgtgga 300  
 acactgcaaa ttattccata gcacggatac caaatctgtg gtttgtccga tatgtgcctc 360  
 gatgccctgg ggagaccca actaccgag cgccaacttc agagagcaca tccagcgccg 420  
 gcaccggttt tcttatgaca cttttgtgga ttatgatgtt gatgaagagg acatgatgaa 480  
 tcagggtgtg cagcgctcca tcatcgacca gtgagcagag tccgtgcttg ctatc 535

<210> 58  
 <211> 479  
 <212> DNA  
 <213> Homo Sapiens

<400> 58  
 gctgaaagaa gcccacatag aactgcttag ggacagcacc actgactcca aagaaaatcc 60  
 cagcagaaag agaaatggaa tgtgcacgga tacacattca ctgctcagta agaggctcaa 120  
 gacatgactg atttgcattt taaagcaaga tgcgatgtcc agagttacag agaattagta 180

## 21

gatgtgtctc atcggttaaat agctctatta tacctctaaa ggtggaattg tcagttttaga	240
ttcataaatg aaaaggtaaa tgagtaatca gaataaacca agtgataatc aaaccatgtc	300
aagattatta gttcagactc tagcctgtta attttcttag ttgatttctg aagctacctg	360
atttattcta ttaaattgta agcttgcaaa ctcaaaataa attggcagat ttacctctca	420
tgttttaatg tgtcaaatta gagagcaaag tataacaggt gccttcactt ttgagactt	479

<210> 59  
 <211> 518  
 <212> DNA  
 <213> Homo Sapiens

<400> 59 gtgccatagt gcaggcttgg ggagctttaa gcctcagtta tataaccac gaaaaacaga	60
gcctcctaga tgtaacattc ctgatcaagg tacaattctt taaaattcac taatgattga	120
ggtccatatt tagtggtact ctgaaattgg tcactttcct attacacgga gtgtgctaaa	180
actaaaaagc attttgaaac atacagaatg ttctattgtc attgggaaat ttttctttct	240
aaccacgtgg aggttagaaa gaagttatat tctggtagca aattaacttt acatcctttt	300
tcctacttgt tatggttgtt tggaccgata agtgtgctta atcctgaggc aaagtagtga	360
atatgtttta tatgttatga agaaaagaat tggtgtaagt ttttgattct actcttatat	420
gctggactgc attcacacat ggcatgaaat aagtcagggt ctttacaat ggtattttga	480
tagatactgg attgtgtttg tgccatattt gtgccatt	518

<210> 60  
 <211> 489  
 <212> DNA  
 <213> Homo Sapiens

<400> 60 gggatgcatt tgtggccatt gttcaaagtg tcaagaacaa gcctctcttc tttgccgaca	60
aactttacaa atccatgaag ggtgctggca cagatgagaa gactctgacc aggatcatgg	120
tatcccgcag tgagattgac ctgctcaaca tccggaggga attcattgag aaatatgaca	180
agtctctcca ccaagccatt gagggtgaca cctccggaga cttcctgaag gccttgctgg	240
ctctctgtgg tggtagaggac tagggccaca gctttggcgg gcacttctgc caagaaatgg	300
ttatcagcac cagccgccat ggccaagcct gattgttcca gctccagaga ctaagggaag	360
ggcaggggtg gggggagggg ttgggttggg ctcttatctt catggagctt aggaaacgct	420
cccactccca cgggccatcg agggccagca cggctgagcg gtgaaaaacc gtagccatag	480
atcctgtcc	489

<210> 61  
 <211> 472  
 <212> DNA  
 <213> Homo Sapiens

<400> 61 atttcaaat ttctgcattc acggagaatg caaatatata gagcacctgg aagcagtaac	60
atgcaaatgt cagcaagaat atttcggtga acggtgtggg gaaaagtcca tgaaaactca	120
cagcatgatt gacagtagtt tatcaaaaat tgcattagca gccatagctg cctttatgtc	180

## 22

```

tgctgtgac ctcacagctg ttgctgttat tacagtccag cttagaagac aatacgtcag      240
gaaatatgaa ggagaagctg aggaacgaaa gaaacttcga caagagaatg gaaatgtaca      300
tgctatagca taactgaaga taaaattaca ggatatcaca ttggagtcac tgccaagtca      360
tagccataaa tgatgagtcg gtctctcttc cagtggatca taagacaatg gacccttttt      420
gttatgatgg ttttaaactt tcaattgtca ctttttatgc tatttctgta ta              472

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<210> 62
<211> 523
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (41)..(41)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (440)..(440)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (442)..(442)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (485)..(486)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (488)..(491)
<223> n is a, c, g, or t

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<220>
<221> misc_feature
<222> (493)..(498)
<223> n is a, c, g, or t

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<400> 62
gacaacagcc ctggagggga acagagtgag agagatgttt ngctctggta cagcctgtgt      60
tgtttgccea gtttctgata tactgtacaa aggcgagaca atacacattc caactatgga      120
gaatggtcct aagctggcaa gccgcattct gagcaaatta actgatatcc agtatggaag      180
agaagagagc gactggacaa ttgtgctatc ctgaatggaa aatagaggat acaatggaaa      240
atagaggata ccaactgtat gctactggga cagactgttg catttgaatt gtgatagatt      300
tctttggcta cctgtgcata atgtagtttg tagtatcaat gtgttacaag agtgattggt      360
tcttcatgcc agagaaaatg aattgcaatc atcaaatggt gtttcataac ttggtagtag      420
taacttacct taccttaccn anaaaaatat taatgtaagc catataacat gggattttcc      480
tcaannannn nannnnnncc ttttgtactt cactcagata cta                          523

```

```

<210> 63
<211> 373
<212> DNA
<213> Homo Sapiens

```

## 23

<400> 63  
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 aactgtgaag atctgacttc gccccccccc cccccatct tcgggaccag gatttgaca 120  
 gaagcacatg cacctaccca tacacccct cttctgagcg tccctgttcc cccatctcgc 180  
 tccctcccag gactctgacc ccagcattct caggcaccag tccctgtccg gaatgccacc 240  
 cacatcttcc atttccatgt cccctcccag agctgggtgga cccaggaac agccactccc 300  
 ctccactctc taccagataa ctgaggaggg gagaggtggg ccgtaacggg cacggatcac 360  
 gatgtaaatt att 373

<210> 64  
 <211> 535  
 <212> DNA  
 <213> Homo Sapiens

<400> 64  
 agcttcagga cgcgtctgca gaggtggagc gactgagaag agaaaaccag gtcttaagcg 60  
 tgagaatcgc ggacaagaag tactacccca gctcccagga ctccagctcc gctgcggcgc 120  
 cccagctgct gattgtgctg ctgggcctca gcgtctgct gcagtgaat cccaggaagc 180  
 tggcacatct tggaaggctc gtccgtctcg gcttttcgct tgaacattcc cttgatctca 240  
 tcagttctga gcggtcatg gggcaacacg gttagcgggg agagcacggg gtagccggag 300  
 aagggcctct ggagcaggctc tggaggggcc atggggcagt cctgggtgtg gggacacagt 360  
 cgggttgacc cagggtgtc tccctccaga gcctccctcc ggacaatgag tccccctct 420  
 tgtctcccac cctgagattg ggcattgggt gcggtgtggg gggcatgtgc tgctgttgt 480  
 tatgggtttt ttttgcgggg ggggttgctt tttctgggg tctttgagct ccaa 535

<210> 65  
 <211> 452  
 <212> DNA  
 <213> Homo Sapiens

<400> 65  
 catgctggac cagatcaact cctgtctgga ccacctggag gagaagaatg accacctcca 60  
 cgccccctc caggagctgc tggagtccaa ccggcagaca cgctggagt tccagcagca 120  
 gctcggggag gccccagtg atgccagccc ctaggctcca agagccccca accgggaccc 180  
 aacctgcct ccttgggcta ggctctggcc tgggcaactc cccctggct tagacacctt 240  
 ctcaagggct ggccttcagg gaccttgggt gggctctgct gcctgggcca cccttctgc 300  
 ctgggcctcc ccttggccta cctgggccag cccccaccac ctggcatgcc ctctggggc 360  
 caagagtggg cctgcaaccc acccacttgc ctgccaccc aactcctggg cgctccccac 420  
 tctgccagg ccttgagtgt ccacattaaa tg 452

<210> 66  
 <211> 323  
 <212> DNA  
 <213> Homo Sapiens

<400> 66  
 cacttaccag tgagcatata tattttaaaa tactttcttt ggatattgta attcttaact 60

## 24

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ggttgtaaata tagaaaagct gggattacat atggtgtgctg gttacagtct aaattttttc 120
atcctcctat gcatcataag catgtttgta atattttcaa aaatagttct actgatgcta 180
caggaatttc aagcctgtgg tgaatgtag tatttaccat agggagtga gtggagttag 240
ggtttcattc aatagagtat tgctgattat acttgagtgg aatcctttcc tcacgiactc 300
ccacagacgt ctgggcctgg aaa 323

```

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<210> 67
<211> 560
<212> DNA
<213> Homo Sapiens

```

```

<400> 67
ggcggaggag aacaaacaga tcatccgcaa acacgcgcag accttcgttg ccctctgtgc 60
cacagatgtg aagttcattt ccaatccgcc ctccatggtg gcagcgggga gcgtggtggc 120
cgcagtgcaa ggcctgaacc tgaggagccc caacaacttc ctgtcctact accgcctcac 180
acgcttcctc tccagagtga tcaagtgtga cccggactgc ctccgggcct gccaggagca 240
gatcgaagcc ctgctggagt caagcctgcg ccaggcccag cagaacatgg accccaaggc 300
cgccgaggag gaggaagagg aggaggagga ggtggacctg gcttgacac ccaccgacgt 360
gcgggacgtg gacatctgag ggcgccaggc agcgggggcg caccgccacc cgcagcgagg 420
gcggagccgg cccaggtgc tccccgaca gtccctcctc tccggagcat tttgatacca 480
gaagggaaag cttcattctc cttgtgtgtg gttgtttttt cctttgtctt ttcccccttc 540
catctctgac ttaagcaaaa 560

```

```

<210> 68
<211> 471
<212> DNA
<213> Homo Sapiens

```

```

<400> 68
gttttgggta tgtttaatct gttatgtact agtgttctgt ttgttattgt tttgttaatt 60
acaccataat gctaatttaa agagactcca aatctcaatg aagccagctc acagtgtgtg 120
gtgccccggt catctagcaa gctgccgaac caaaagaatt tgcacccgc tcggggccca 180
cgtggttggg gccctgccct ggcagggtca tctgtgtctc ggaggccatc tcgggcacag 240
gccacccccg cccacccct ccagaacacg gctcacgctt acctcaacca tcttggtgc 300
ggcgtctgtc tgaaccacgc gggggccttg agggacgctt tgtctgtcgt gatggggcaa 360
gggcacaagt cctggatggt gtgtgtatcg agaggccaaa ggctggtggc aagtgcacgg 420
ggcacagcgg agtctgtcct gtgacgcgca agtctgaggg tctgggcggc g 471

```

```

<210> 69
<211> 518
<212> DNA
<213> Homo Sapiens

```

```

<400> 69
aattcctgcc attctgggga ttcttgagg aattcttgct ttgctaattc tgattctgct 60
gctcttgctg tttcttcgga ggagagcggg ggtcaaagag cccttactgc cccagagga 120

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## 25

tgacacccgg gacaacgttt attactatga tgaagaagga ggcggagaag aggaccagga 180  
 ctttgacttg agccagctgc acaggggcct ggacgctcgg cctgaagtga ctgtaacga 240  
 cgttgacca accctcatga gtgtcccccg gtatcttccc cgccctgcc atcccgatga 300  
 aattggaaat ttattgatg aaaatctgaa agcggctgat actgaccca cagccccgcc 360  
 ttatgattct ctgctcgtgt ttgactatga aggaagcggg tccgaagctg ctagtctgag 420  
 ctccctgaac tcctcagagt cagacaaaga ccaggactat gactacttga acgaatgggg 480  
 caatccgttc aagaagctgg ctgacatgta cggaggcg 518

<210> 70  
 <211> 182  
 <212> DNA  
 <213> Homo Sapiens

<400> 70  
 cttttcactg tgttgaggtt ttctggagtg agcactcacg ccctaagcgc acattcatgt 60  
 gggcatttct tgcgagcctc gcagcctccg gaagctgtcg acttcatgac aagcattttg 120  
 tgaactaggg aagctcaggg gggttactgg cttctcttga gtcacactgc tagcaaatgg 180  
 ca 182

<210> 71  
 <211> 538  
 <212> DNA  
 <213> Homo Sapiens

<400> 71  
 tgaggagcca gcgtctaggg cagcagccgc ttctagaag accaggatcat gatgatgggc 60  
 agcgcgccgag tggcggagct gctgctgctc cagggcgagg agcccaactg cgccgacccc 120  
 gccactctca cctgacccgt gcacgacgct gcccgggagg gcttccctgga cagctgggtg 180  
 gtgctgcacc gggccggggc gcggctggac gtgcgcgatg cctggggccg tctgcccgtg 240  
 gacctggctg aggagctggg ccacgcgatg gtgcacgggt acctgcgcgc ggctgcgggg 300  
 ggcaccagag gcagtaacca tgcccgcata gatgccacgg aaggctccctc agacatcccc 360  
 gattgaaaga accagagagg ctctgagaaa cctcgggaaa cttagatcat cagtcaccga 420  
 aggtcctaca gggccacaac tgccccgcc acaaccacc ccgcttctgt agttttcatt 480  
 tagaaaatag agctttttaa aatgtcctgc cttttaacgt agatatatgc cttcccc 538

<210> 72  
 <211> 513  
 <212> DNA  
 <213> Homo Sapiens

<400> 72  
 atattagtta ccctgggtgtg ctgtattctc taaaaccttt aaatgtttgc atgcagccat 60  
 tcgtcaaatg tcaaatattc tctctttggc tggaatgaca aaaactcaaa taaatgtatg 120  
 attaggagga catcataacc tatgaatgat ggaagtccaa aatgatggta actgacagta 180  
 gtgttaatgc cttatgttta gtcaaaactct catcttaggtg acagcctggt gactccagaa 240  
 tggagccagt catgctaaat gccatatact cacactgaaa catgaggaag caggtagatc 300  
 ccagaacaga caaaactttc ctaaaaacat gagagtccag gctgtctgag tcagcacagt 360

## 26

aagaaagtcc tttctgcttt aactccttaga aaaaagtaat atgaagtatt ctgaaattaa 420  
 ccaatcagtt tatttaaate aatttatatta tattcttctg ttccctggatt cccattttac 480  
 aaaaccact gttctactgt tgtattgccc agt 513

<210> 73  
 <211> 530  
 <212> DNA  
 <213> Homo Sapiens

<400> 73  
 ggatttgtgt tcttacagta cttgaaaata ttaaggaag agatgaagct ctgcagtttt 60  
 ttctatgtgg gatgattact tttttaagga ggattaattc tgaggtagta tagtaactaa 120  
 aggggaatat atgaattggt taacaaatta gaatttggtt acaactactt gaatttttaa 180  
 attatgtcaa aacttacatt acttgccaag cagtatgatg ttataggaaa cataaataag 240  
 attacagagg tatcaatttg gttaaaattc accattttat aagactaagc aataatctta 300  
 acaacctctt tctgaatat ttaaatgtgt ttgtatgggt ttatgactaa ttgttactga 360  
 tttagagact aagccctctt aaaaccttta gttaaatata aaaagaaatt atatatatct 420  
 tgcctccctg atggaaaact atataaaatt gtagacttaa aaggtttggt gaaatacatt 480  
 aggatatcag aaaactaaat atatggaggt gctttatgac tattacatgt 530

<210> 74  
 <211> 406  
 <212> DNA  
 <213> Homo Sapiens

<400> 74  
 ggctgcctgc ggatgaagga ccagtgtgac aagtgcggg agatcttgtc tgtggactgt 60  
 tccaccaaca accctccca ggctaagctg cggcgggagc tcgacgaatc cctccaggtc 120  
 gctgagaggt tgaccaggaa atataacgag ctgctaaagt cctaccagtg gaagatgtc 180  
 aacacctcct ccttgctgga gcagctgaac gagcagttta actgggtgtc cgggctggca 240  
 aacctcacgc aaggcgaaga ccagtactat ctgcgggtca ccacgggtgc tccccacact 300  
 tctgactcgg acgttccttc cgggtgcact gaggtggctg tgaagctctt tgactctgat 360  
 cccatcactg tgacggtccc tgtagaagtc tccaggaaga acccta 406

<210> 75  
 <211> 286  
 <212> DNA  
 <213> Homo Sapiens

<400> 75  
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 accagtacta tctgcgggtc accacgggtg cttccacac ttctgactcg gacgttcctt 120  
 ccgggtgcac tgagggtgtc gtgaagctct ttgactctga tccatcact gtgacggtcc 180  
 ctgtagaagt ctccaggaag aaccctaaat ttatggagac cgtggcggag aaagcgtgc 240  
 aggaataccg caaaaagcac cgggaggagt gagatgtgga tgttgc 286

<210> 76



<211> 436  
 <212> DNA  
 <213> Homo Sapiens

<400> 76  
 gaaagactgt gctgtccttt aacatagggt tttaaagact aggatattga atgtgaaaca 60  
 tccgttttca ttgttcactt ctaaaccaaa aattatgtgt tgccaaaacc aaaccagggt 120  
 tcatgaatat ggtgtctatt atagtgaac atgtactttg agcttattgt ttttattctg 180  
 tattaaatat tttcagggtt ttaaacacta atcacaaact gaatgacttg acttcaaaag 240  
 caacaacctt aaaggccgtc atttcattag tattcctcat tctgcacccg ggcttgaaaa 300  
 acagctctgt tgaatcacag tatcagtatt ttcacacgta agcacattcg ggccatttcc 360  
 gtggtttctc atgagctgtg ttcacagacc tcagcagggc atcgcattga ccgcaggagg 420  
 gcagattcgg accact 436

<210> 77  
 <211> 429  
 <212> DNA  
 <213> Homo Sapiens

<400> 77  
 tcggctactc ttttgtgatg cacaccagcg ctggtgcaga aggctctggc caagccctgg 60  
 cgtcccccg ctcctgcctg gaggagtta gaagtgcgcc attcatcgag tgtcacggcc 120  
 gtgggacctg caattactac gcaaacgctt acagcttttg gctcgccacc atagagagga 180  
 gcgagatgtt caagaagcct acgccgtcca cctgaaggc aggggagctg cgcacgcacg 240  
 tcagccgctg ccaagtctgt atgagaagaa cataatgaag cctgactcag ctaatgtcac 300  
 aacatggtgc tactttctct tctttttgtt aacagcaacg aaccctagaa atatatcctg 360  
 tgtacctcag tgtccaatat gaaaaccgta aagtgcctta taggaatttg cgtaactaac 420  
 acaccctgc 429

<210> 78  
 <211> 195  
 <212> DNA  
 <213> Homo Sapiens

<400> 78  
 tccccctgta gactagtgcc gtgggagtag ctgctgcccc gctgctgtgg cccctccgt 60  
 gatccatcca tctccaggga gcaagacaga gacgcaggat ggaaagcgga gttcctaaca 120  
 ggatgaaagt tcccccatca gtccccccag tacctccaag caagtagctt tccacatttg 180  
 tcacagaaat cagag 195

<210> 79  
 <211> 301  
 <212> DNA  
 <213> Homo Sapiens

<400> 79  
 tgggtgtggg agccctttgg agaacgccag tctccaggtc cccctgcacg tatcgagttt 60  
 gcaatgtcac aacctctctg atcttgtgct cagcatgatt ctttaataga agttttatct 120  
 ttcgtgcact ctgctaata tgtgggtgag ccagtggaa acgaggagcc tgtgctggtt 180

## 28

tgcagattgc ctctaatga cgcggctcaa aaggaaacca agtggtcagg agttgtttct 240  
gacccactga tctctactac cacaaggaaa atagtttagg agaaaccagc ttttactgtt 300  
t 301

<210> 80  
<211> 459  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> misc\_feature  
<222> (164)..(164)  
<223> n is a, c, g, or t

<400> 80  
ggaaacgttc ccagttcatt ttcagtcctg ttgtgagcac agttctgaag ggtttattat 60  
tgtcaaaaata agttttgttt tgtttgttt atgttgggtt tttaatgttg tctcttgacc 120  
cttaatgctc aggttcttgt gggagttaat cagccacatc caangttacc ttgaggggga 180  
agaagagggt gatgctcaga agctaaacaa gacaggggcc acatgaccct ctattgatta 240  
gccccaaagta gaaagtcctg tggttttatg tttaatggta atagttgatc atatatggca 300  
taattttcta tcagcttcct actcagtcac tataaacaca gacttgaaat agtactttaa 360  
atgtccaaat acctaaatgt gctaaactgg aggtaactat ttctaggtag ttgaattttt 420  
gaaagtcatg atcagccaca caactgtttt gtacatact 459

<210> 81  
<211> 394  
<212> DNA  
<213> Homo Sapiens

<400> 81  
aatccttatt gttcagagtt gtttgggggt tctgtttcag agcataaaac ctaaaggtta 60  
tagtagaaca aggcaccttc ttaaaagaaa tcttgcttca gaccatcagt tacagagaat 120  
ttcctaaagt aaaattgaag caactacaac ttctccttag acactttgga atctaaccac 180  
ttaaggacct ttttaaagag atagcttctc ttctttctga agatcaattt ctccaaggc 240  
caagattgtc cttttctccc atttcttctc agctattgca aatgagggaa gaacattatt 300  
catctctcct cccctttttt ttctgattct ttttccagtc agttttgtc ctgggttcaa 360  
gtagtattac caccctttca caagcaacag actc 394

<210> 82  
<211> 514  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> misc\_feature  
<222> (89)..(89)  
<223> n is a, c, g, or t

<400> 82  
gtcactaca ctattcattg cacacaaatg aatttttcac tttttaagat gcattcttgg 60  
tgctcaaacc agatcgaagt ttgtctctna aagctattgt ctgcacaggc tgctgcatgc 120

## 29

tctgttggtta aatggatgga caggctattc taaattttgg ttgatacttt tgctactatg 180  
ggcaattaac ttgaaaaaaa taatcgatcc caactctgtg ctctgatgta cctcttctgc 240  
cccttttatg acacctttga ccaaatgcct tctatgggtc acagtgcagg cacaaaacta 300  
cctctgatac agaaggttct ttacaagctt attttacata ccgtgaatcc ctcacctaaa 360  
gggagagggtg aaagcaaaga ctgctttgaa tgggtattga gggagattgt gtccatacca 420  
agccaccctg aagaagtatt tcaactgcag tagaactgtg gatttgtgct gtcatttcac 480  
cttggaaataa acacctatct ctaagcagga ccaa 514

<210> 83  
<211> 299  
<212> DNA  
<213> Homo Sapiens

<400> 83  
caccaaatta cctaggctga ggtagagag attggccagc aaaaactgtg ggaagatgaa 60  
ctttgtcatt atgatttcat taccacatga ttatagaagg ctgtcttagt gcaaaaaaca 120  
tacttacatt tcagacatat ccaaaggga taccacatt ttgttaagaa gttgaactat 180  
gactggagta aaccatgtat tcccttatct ttactttttt ttctgtgaca tttatgtctc 240  
atgtaatttg cattactctg gtggattgtt ctagtactgt attgggcttc ttctgtaat 299

<210> 84  
<211> 219  
<212> DNA  
<213> Homo Sapiens

<400> 84  
ttatcgccct gagaagatct accccagga gaatctgaga catcttgcct acttttcttt 60  
attagcttct tctcatcca tttcttttat accttctctt tttggggagt tggtatgcca 120  
tgatttttgg tatttatgta aaaggattat tactaattct atttctctat gtttattcta 180  
gttaaggaaa tggtgagggc aagccaccaa attacctag 219

<210> 85  
<211> 518  
<212> DNA  
<213> Homo Sapiens

<220>  
<221> misc\_feature  
<222> (61)..(65)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (71)..(71)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (73)..(73)  
<223> n is a, c, g, or t

<220>  
<221> misc\_feature  
<222> (112)..(112)

<223> n is a, c, g, or t

<220>

<221> misc\_feature

<222> (163)..(163)

<223> n is a, c, g, or t

<220>

<221> misc\_feature

<222> (295)..(295)

<223> n is a, c, g, or t

<400> 85

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aaggactggt atctttctgt gagcaataag gactggataa agactgcata tccttggtgc      60
nnnnncagca ncnatacaat aaggagggtt ttaatgtgaa gcaggcaatc tnccagcccc      120
ttctggtctt ggatgaaata gttgcacaga gtattgcacc aanaatacac aatggagggt      180
gaaaagttca acatatttta agtcaattaa tcaaattgca ttgattcttg atgctttctt      240
agaggcctac atgatttctt agattgctct gataaactat cataaggggt ccacntcccc      300
tcatttagct cccccagga tttcttttcc cccatgtcat acaccagtc ctaaatcaac      360
ccccagggt atccttccat cccttctgca gaggaactt ttgtcagact ctgcaacaaa      420
ctcctagctc tatccagagt gtccctctgt gctaagattg gtatctttct cctcaaaagc      480
ctggatggtg aatgggggtg cattagtcag aattctcc                               518

```

<210> 86

<211> 458

<212> DNA

<213> Homo Sapiens

<400> 86

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taaaaaacctg tatctgaccc actttgtaat ttttgtcca atatccattc tgtagacttt      60
tgaaaaaaa gtttttaatt tgatgcccaa tatattctga ccgttaaaaa attcttggtc      120
atatgggaga agggggagta atgacttgta caaacagtat ttctggtgta tattttaatg      180
tttttaaaaa gagtaatttc atttaaatat ctgttattca aatttgatga tgttaaatgt      240
aatataatgt attttctttt tattttgac tctgtaattg cactttttaa gtttgaagag      300
ccattttggt aaacggtttt tattaaagat gctatggaac ataaagtgt attgcatgca      360
atttaaagta acttatttga ctatgaatat tatcggatta ctgaattgta tcaatttgtt      420
tgtgttcaat atcagctttg ataattgtgt accttaag                               458

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<210> 87

<211> 336

<212> DNA

<213> Homo Sapiens

<400> 87

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gggatcctat ttagctctta gtaccactaa tcaaaagttc ggcatgtagc tcatgatcta      60
tgctgtttct atgtcgtgga agcaccggat ggggtagtg agcaaatctg ccctgctcag      120
cagtcaccat agcagctgac tgaaaatcag cactgcctga gtagttttga tcagtttaac      180
ttgaatcact aactgactga aaattgaatg ggcaaataag tgcttttgct tccagagtat      240
gcgggagacc ctccacctc aagatggata tttcttcccc aaggatttca agatgaattg      300
aaatttttaa tcaagatagt gtgctttatt ctgttg                               336

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## 31

<210> 88  
 <211> 521  
 <212> DNA  
 <213> Homo Sapiens

<400> 88  
 atatcttctt caggctctga caggcctcct ggaaacttcc acatatTTTT caactgcagt 60  
 ataaagtcag aaaataaagt taacataact ttcactaaca cacacatatg tagattttcac 120  
 aaaatccacc tataattggt caaagtgggt gagaatatat tttttagtaa ttgcatgcaa 180  
 aattttttcta gcttccatcc tttctccctc gtttcttctt tttttggggg agctggtaac 240  
 tgatgaaatc ttttccacc ttttctcttc aggaaatata agtgggtttg tttggttaac 300  
 gtgatacatt ctgtatgaat gaaacattgg agggaaacat ctactgaatt tctgtaattt 360  
 aaaatatttt gctgctagtt aactatgaac agatagaaga atcttacaga tgctgctata 420  
 aataagtaga aaatataaat ttcataccta aaatatgcta ttttaaaatc tatttcctat 480  
 attgtatttc taatcagatg tattactctt attatttcta t 521

<210> 89  
 <211> 503  
 <212> DNA  
 <213> Homo Sapiens

<400> 89  
 gtggctatcc actgttagtt cagaagctgg gcttggacta ctcttatgat ttagctccac 60  
 gagccaaaat tttccggcgt gaccaaggga aagtgactga tacggcatcc atgaaatata 120  
 tcatgcgata caacaattat aagaaggatc cttacagtag aggtgacccc tgtaatacca 180  
 tctgctgccg tgaggacctg aactcaccta acccaagtcc tggaggttgt tatgacacaa 240  
 aggtggcaga tatctaccta gcatctcagt acacatccta tgccataagt ggtccacag 300  
 tacaagggtg cctccctgtt tttcgtggg accgtttcaa caaaactcta catcagggca 360  
 tgccagaggt ctacaacttt gatatttatta ccatgaaacc aattttgaaa cttgatataa 420  
 aatgaaggag ggagatgacg gactagaaga ctgtaaataa gataccaaag gcactatttt 480  
 agctatgttt ttcccatcag aat 503

<210> 90  
 <211> 275  
 <212> DNA  
 <213> Homo Sapiens

<400> 90  
 ccccatcacg gaggggtccag actgtccact cgggggtgga gtgagactga ctgcaagccc 60  
 caccctcctt gagactggag ctgagcgtct gcatacgaga gacttggttg aaacttggtt 120  
 ggtccttgte tgcaccctcg acaagaccac actttgggac ttgggagctg gggctgaagt 180  
 tgctctgtac ccatgaactc ccagtttgcg aattaataag agacaatcta ttttggtact 240  
 tgcacttggt attcgaacca ctgagagcga gatgg 275

<210> 91  
 <211> 405  
 <212> DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 91

tcattctgatg	tttctatagt	cacttttgcca	gctcaaaaga	aaacaatacc	ctatgtagtt	60
gtggaagttt	atgctaatat	tgtgtaactg	atattaaacc	taaatgttct	gcctaccctg	120
ttggtataaa	gatatattga	gcagactgta	aacaagaaaa	aaaaaatcat	gcattcttag	180
caaaattgcc	tagtatgtta	atttgctcaa	aatacaatgt	ttgatattat	gcactttgtc	240
gctattaaca	tccttttttt	catgtagatt	tcaataattg	agtaatttta	gaagcattat	300
tttaggaata	tatagttgtc	acagtaaata	tcttgttttt	tctatgtaca	ttgtacaaat	360
ttttcattcc	ttttgctctt	tgtggttgga	tctaacacta	actgt		405

&lt;210&gt; 92

&lt;211&gt; 375

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 92

aagctatgtg	tatcttctgt	gtaaagcagt	ggcttcactg	gaaaaatggg	gtggctagca	60
tttccctttg	agtcatgatg	acagatgggt	tgaaaaccat	ctaagtttgc	ttttgaccat	120
cacctcccag	tagcaatttg	ctttcataat	ccatttagca	atccaggcct	ctgttgaaaa	180
gataatatga	gggagaaggg	aacacatttc	cttctgaact	tacttcoccta	agtcactttc	240
cttatgtatc	atctaataca	atgatgggtg	agtgaaaata	cagaaggggt	gtttgagtat	300
tcagatttca	taaaacactt	ccttggaata	tagctgcatt	aacttggaag	gaagcctggt	360
gggccagaag	acaga					375

&lt;210&gt; 93

&lt;211&gt; 533

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 93

gctggtgtgt	gtgtcaaacc	ctcactcacc	cacgcactca	cacacagcat	tctgttctcc	60
atgcaaagtt	aagatcgaat	ccatccgctt	gtaggggaaa	aaaaggaaaa	aaattaacca	120
gagagggtct	gtaatctcgc	agagcacagg	cagaatcggt	ccttccttgc	tgcatctcct	180
ccttagacta	atagacgttt	tggaagttc	ggctagtgtt	cgtgtgtttg	tcgtagcacc	240
cagagcctcc	accaaaccct	ctccatgtct	ttacctccca	gtcgcctetaa	gatctgcttg	300
aagtctcgta	tttgtaactgc	tttctgcttt	tctcccaccc	ctcctagcac	ccccacatcc	360
cccatctagt	aacatctcag	aaatttcac	cagaggaaca	aaaaaattaa	aaatagaaca	420
tagcaaagca	aagacagaat	gccccccccc	aaatatgttc	ctgtccctgt	ctgggagttg	480
tgttatttaa	agatattctg	tatgttgtat	cttttgcattg	tagcttcctt	aat	533

&lt;210&gt; 94

&lt;211&gt; 413

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 94

atctggaagg	ctctgatcca	cctgagcgac	ctccgggagt	acaggcgctt	tgagaaggag	60
------------	------------	------------	------------	------------	------------	----

## 33

aagctcaagt ccagtgga caatgataat ccccttttca agagcgccac cagcagggtc 120  
 atgaacccca agtttctga gagttaggag cacttggtga agacaaggcc gtcaggaccc 180  
 accatgtctg ccccatcacg cggccgagac atggcttggc cacagctctt gaggatgtca 240  
 ccaattaacc agaaatccag ttattttccg ccctcaaaat gacagccatg gccggccgggt 300  
 gcttctgggg gctcgtcggt gggacagctc cactctgact ggacagctct ttgcatggag 360  
 acttgaggag ggcttgaggt tggtagggt aggtgcgtgt ttctgtgca agt 413

<210> 95  
 <211> 465  
 <212> DNA  
 <213> Homo Sapiens

<400> 95  
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 ccacccgagg gccagtgagt cggggaagag gacttctcac tcccagagca agaggagtcc 120  
 cccaactgg gtacagacct ccaccgccac ccccgacaca agagacttat ggagaatatg 180  
 actatgatga tggatatggc actgcttatg atgaacagag ttatgattcc tatgataaca 240  
 gctatagcac ccagcccaa agtgggtctg attactatga ttacggacat ggactcagtg 300  
 aggagactta tgattctcac gggcaagaag agtggactaa ctcaagacac aaggcacctt 360  
 cagcgaggac agcaaagggc gtctacagag accagccata tggcagatac tgattgtact 420  
 gtctgatgtt gtgaaatagc caatctccac cagtctgtga tactg 465

<210> 96  
 <211> 537  
 <212> DNA  
 <213> Homo Sapiens

<400> 96  
 gagaacacgg tggcagagac ggagtgccgc tatgccctgc agctgcagca gatccagggg 60  
 ctcatcagca gcacgagggc ccagctgagc gagctccgca gtgagatgga gtgccagaac 120  
 caagagtaca agatgctgct ggacatcaag acacgtctgg agcaggagat cgccacctac 180  
 cgcagcctgc tcgagggcca ggacgccaag aagcgtcagc ccccgtagca cctctgttac 240  
 cagcacttct agtgcctctg ttaccaccac ctctaagtc tctggtcgcc gcaacttctga 300  
 tgtccgtagg ccttaaactc gcctggcgtc ccctccctct gtcttcagca ccagaggag 360  
 gagagagccg gcagttccct gcaggagaga ggaggggctg ctggacccaa ggctcagtc 420  
 ctctgtctc aggacccctc gtctgactc tctctgatg gtgggccctc tgtgctcttc 480  
 tctccggtc ggatctctct cctctctgac ctggatacgc tttggtttct caacttc 537

<210> 97  
 <211> 372  
 <212> DNA  
 <213> Homo Sapiens

<400> 97  
 aactttaact tagagcttca ttactttaag aatggaaaac aacctctgag ttgatttcc 60  
 caaagtttca taaagccctc aagctcatga ttttcatcaa ctctttgccc acatagtcac 120  
 ttacctccac agccgtttgt tgtcatagaa ggggtgggtg tgtttggatt tgattttttt 180

## 34

caacttgcag tgagaaatag gataggtgac aaaaccttac ttgttttctt aagacaattc 240  
 agtgcttgag catctctgtc agaaatggaa tgaaatactg ttagccaatt agaattatct 300  
 tatgtattgt tattgtgttt tgctgatttt tatatgaaaa tataattatt cattcttgat 360  
 ctctggaagc aa 372

<210> 98  
 <211> 365  
 <212> DNA  
 <213> Homo Sapiens

<400> 98  
 gggagccaag gctttatacg tctaaagaaa atattcagta gctgaatccg cccagtgata 60  
 gcctgtgggc accagcagca agggctgcca tgggatacag caccatctc caaagacctc 120  
 tattacataa aactgcttc ttacaggaaa caaacctctt ctgggatctc cttttgtgaa 180  
 aaccagtttg atgtgctaaa agtaaaaagt ctattttcca gtgtggtctt gttcagaagc 240  
 agccagattt ccaatgttgt ttttccctc cactcagaaa cccctgccct ttcccttcag 300  
 aaaacgatgg caggcattcc tctgagttta caagcagaga ctcactccaa cccaaactag 360  
 ctggg 365

<210> 99  
 <211> 465  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> misc\_feature  
 <222> (110)..(110)  
 <223> n is a, c, g, or t

<400> 99  
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 ccttaaaagg gagtcttatg ttttcaacta cagatagttg taagggatcn tacagaagat 120  
 attgatgata gttgaaatat tcttagaagg ggtgtgtatg tctagctgtg tctaccatgt 180  
 gtatgtattc ttgacaagca gtataaaata cctgtgattt ttctttacat tagggataat 240  
 gcataaggaa ttaatcttca tatatattat catccctaatt gtagcagggg gaagtattta 300  
 attgcccatg atatgtatct tactttact atgccagaga ggaaactata aagtaattac 360  
 acatgtaatc ttgggttttt cacatatgta ggtattcatt ttgagtaggt tgaagaagaa 420  
 aaaaaatatt taaatgaatt gaattcctga tgggtagta tcaat 465

<210> 100  
 <211> 515  
 <212> DNA  
 <213> Homo Sapiens

<400> 100  
 gaactctgca tcttcatggt ttacagaaat tgggtgcaggc agccagcagt tagattccat 60  
 tcatgtaaca cagttggaga gagataccgt tttagtgtgt ttagacaaat ttgtgaaaat 120  
 tgtaaactca caaggaaaat taaaatcaag taagaaactg gcctctgagt taagttttga 180



## 35

ttttcgcatt gaatctgtag tatgccttca agacagtgtg ttggctttct ggaaacatgg 240  
 gatgcagggt aaaagcttca agtcagatga ggttaccag gagatttcag atgaaacaag 300  
 agttttccgc ttattaggat cagacagggt tgtcgttttg gaaagtaggc caacagaaaa 360  
 tcctactgca cacagcaatc tctacatctt ggctggacat gaaaatagtt actaagcaac 420  
 agaaactgat ctcaaatgac aggaaaatga atatactcca ttgaaaggga aaataaggaa 480  
 attcaataca aactgcacta tgatttgcct taact 515

<210> 101  
 <211> 525  
 <212> DNA  
 <213> Homo Sapiens

<400> 101  
 ctcagagcca cccctaaaga gatcctttga tattttcaac gcagccctgc tttgggctgc 60  
 cctgggtgctg ccacacttca ggctcttctc ctttcacaac cttctgtggc tcacagaacc 120  
 cttggagcca atggagactg tctcaagagg gcactgggtg cccgacagcc tggcacaggg 180  
 cagtgggaca gggcatggcc aggtggccac tccagacccc tggcttttca ctgctggctg 240  
 ccttagaacc tttcttacat tagcagtttg ctttgtatgc actttgtttt tttctttggg 300  
 tcttggtttt tttttccact tagaaattgc atttcctgac agaaggactc aggttgtctg 360  
 aagtcactgc acagtgcac tcagcccaca tagtgatggt tccccgttc actctactta 420  
 gcatgtccct accgagtctc ttctccactg gatggaggaa aaccaagccg tggcttcccg 480  
 ctcagccctc cctgcccctc ccttcaacca tccccatgg gaaat 525

<210> 102  
 <211> 418  
 <212> DNA  
 <213> Homo Sapiens

<400> 102  
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 gaacacacaa cgtcttggag gcgcagagga ggaacgagct aaaacggagc ttttttgccc 120  
 tgctgaccca gatccccgag ttggaaaaca atgaaaaggc cccaaggta gttatcctta 180  
 aaaaagccac agcatacatc ctgtccgtcc aagcagagga gaaaagctc atttctgaag 240  
 aggacttggt gcggaaacga cgagaacagt tgaaacacaa acttgaacag ctacggaact 300  
 cttgtgcgta aggaaaagta aggaaaacga ttccttctaa cagaaatgtc ctgagcaatc 360  
 acctatgaac ttgtttcaaa tgcgatgatca aatgcaacct cacaaccttg gctgagtc 418

<210> 103  
 <211> 462  
 <212> DNA  
 <213> Homo Sapiens

<400> 103  
 aacatccgcc tggtaccag tcgtcttggc tgggcacttc caccgcacc tcattcctac 60  
 atcaatgagt ggctccaaat agacctgggg gaggagaaga tcgtgagggg catcatcatt 120  
 cagggtggga agcaccgaga gaacaagggt ttcattgagga agttcaagat cgggtacagc 180  
 aacaacggct cggactggaa gatgatcatg gatgacagca aacgcaaggc gaagtctttt 240

## 36

```

gagggcaaca acaactatga tacacctgag ctgcggaactt ttccagctct ctccacgcga      300
ttcatcagga tctaccccga gagagccact catggcggac tggggctcag aatggagctg      360
ctgggctgtg aagtggaaagc ccctacagct ggaccgacca ctccaacgg gaacttggtg      420
gatgaatgtg atgacgacca ggccaactgc cacagtggaa ca                          462

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<210> 104
<211> 370
<212> DNA
<213> Homo Sapiens

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<220>
<221> misc_feature
<222> (168)..(168)
<223> n is a, c, g, or t

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<400> 104
gcaaatatct taccaggcag cctatgaatt aacccaaaga agctttgggtt ggttttggtg      60
gatttttatc atgccatggtt ggacatgaga ttttttagat cttccttccc acattgctag      120
acgtctcact caaagacatt tgttgggagt cacatttgca tcataganga gacagtccat      180
tcacttagt taaattggat tgagaatgcc ttttgtttcc aggaaaatat tgatcaccat      240
gaaagaagaa tagttttttg tccccagaga cattcattta gttgatataa tctaccaga      300
aggaaagcac taagaaacac tcgtttgttg tttttaagg caacagactt aaagttgtcc      360
tcagccaagg                                     370

```

```

<210> 105
<211> 434
<212> DNA
<213> Homo Sapiens

```

```

<400> 105
caggtgtatc tgcacagtgg tcgccccaca gcagaccatg tgttcacggg atgcccgcac      60
aaaacagctg aggcagctac tggagaaggt gcagaacatg tctcaatcca tagaggctctt      120
ggacaggcgg acccagagag acttgcagta cgtggagaag atggagaacc aaatgaaagg      180
actggagtcc aagttaaac aggtggagga gagtcataag caacacctgg ccaggcagtt      240
taagggctaa cttaaaagag ttttttcaat gctgcagtga ctgaagaagc agtccactcc      300
catgtaacca tgaaagagag ccagagagct ttttgacca tgcattttta ctattatatt      360
ccaatactta gcaccatttc actaaggaac cttgaatata accaggatcc tcctttgcat      420
gcgactgtag ctgc                                  434

```

```

<210> 106
<211> 503
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> misc_feature
<222> (158)..(158)
<223> n is a, c, g, or t

```

```

<220>

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<221> misc\_feature  
 <222> (216)..(217)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (231)..(231)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (250)..(250)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (261)..(261)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (291)..(291)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (297)..(297)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (341)..(341)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (352)..(352)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (365)..(365)  
 <223> n is a, c, g, or t

<400> 106  
 gcgggccaca gacgtcggaa gaaactcccg tatttgcagc tggaactgca gccacggcg 60  
 ccccggtttt cctccccgcc ctgtccctct ctggtcaaac aacatactaa agaggcgagg 120  
 caatgactgt tggccagttc tcaccgggga aaaaccnnc tgttaggatg gcatgaacat 180  
 ttccttagat cgtggtcagc tccgaggaat gtggcncca ggctctttga ngagccatgg 240  
 gctgcacccn ggccgtaggg ntagtgtaac tcgcatccca ttgcagtgcc ngtttctttg 300  
 actgtgttgc tgtctcttag attaacctg ctgaggctcc nacatagctc cntggacctg 360  
 tgtcntagta catactgaag cgatgggtcag agtgtgtaga gtgaagttgc tgtgccaca 420  
 ttgtttgaac tcgcgtaccc cgtagataca ttgtgcaacg ttcttctgtt attcccttga 480  
 ggtggttaact tcgtatgttc agt 503

<210> 107  
 <211> 556  
 <212> DNA  
 <213> Homo Sapiens

<400> 107  
 ggagacttga gcttgacctt aggatatgca ttaaccactc tacagaactc caetcagtac 60

## 38

tgtacaggggt ggctgtgggc ctagaagttc agtttttact gaggaatat ttccattaac 120  
 agcaattatt atattgaagg ctttaataaa ggccacagga gacattacta tagcatagat 180  
 tgtcaaagt aaatttactg agcgtgtttt ataaaaaact cacagggtgt tgaggccaaa 240  
 acagatttta gacttacctt gaacggataa gaatctatag ttcactgaca cagtaaaatt 300  
 aactctgtgg gtggggggcg ggggcatagc tctaacttaa tatataaaat gtgtgatgaa 360  
 tcaacaagat ttccacaatt cttctgtcaa gcttactaca gtgaaagaat gggattggca 420  
 agtaacttct gacttactgt cagttgtact tctgtccat agacatcagt attctgcat 480  
 catttttgat gactacctca gaacataaaa aggaacgtat atcacataat tccagtcaca 540  
 gtttttggtt cctctt 556

<210> 108  
 <211> 543  
 <212> DNA  
 <213> Homo Sapiens

<400> 108  
 ctgacctctt tgaagttgca gaatgctttg aaattctaát ggtatctgaa atatcagctc 60  
 atagaaagta acaaaatttg ctgtcacctt aaataagaca ttttaatttt gttataatgt 120  
 acaatttaga agtttgatta attatattat ctatttaggc attaatataa aagaggtagg 180  
 agtctgttat ttaaaaaaag cattaaattt aaaaaaaaac tgtcttgtct acttttagct 240  
 tcattctccc atattttgaa ggggtgtgtaa cttcagctct gcaggattgc atggggtaaa 300  
 acttgttacc aacacatgtg aaccattgct acattgtagg ttgtgatcat tttgccccac 360  
 tgaagcccat gtatctgacc ttacgtgctt tttgaactag gagaatcggg ctaatttatt 420  
 aatgatgata attataatgt atctgtacag cactttttac atttggaag tgctttccaa 480  
 tccatgtagg ttactagtta ttacagctgt aaggataaaa cacgtcatgt ggattcattt 540  
 tga 543

<210> 109  
 <211> 458  
 <212> DNA  
 <213> Homo Sapiens

<400> 109  
 agaaaaattg ccaatcttct ctactttcta tttttatgat gacaatcaaa gccggcctga 60  
 gaaacactat ttgtgacttt ttaaacgatt agtgatgtcc ttaaaatgtg gtctgccaat 120  
 ctgtacaaaa tggctctatt tttgtgaaga gggacataag ataaaatgat gttatacatc 180  
 aatatgtata tatgtatttc tatatagact tggagaatac tgccaaaaca tttatgacaa 240  
 gctgtatcac tgcttcgtt tatatttttt taactgtgat aatccccaca ggcacattaa 300  
 ctgttgcaact tttgaatgtc caaaatttat attttagaaa taataaaaag aaagatactt 360  
 acatgttccc aaaacaatgg tgygtgaat gtgtgagaaa aactaacttg atagggtcta 420  
 ccaatacaaa atgtattacg aatgcccctg ttcatggt 458

<210> 110  
 <211> 412  
 <212> DNA

## 39

&lt;213&gt; Homo Sapiens

&lt;400&gt; 110

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gtcaaaccat gactcgaca tggcaaaaga acggggccac agtacagcct cacattcttc      60
ttccaattct gaagatacag agatgtgatg aaaacaagta atagctttgg ctgtttatatt    120
gatagctggt tctgggtatt taataggaat cctttctcaa ggaatgagtt gtgacctgtt    180
tactgtctct ttagaagaaa aactccactg gaaaccattc accatgtgtg actgtcttct    240
gttatcatth gtcttacagg cggctattgc agacggctaa tttatgctta acttaggaag    300
agataaggca agagctagat ttttttcatg tgatcttttc caagcttcaa cttaacttaa    360
ctacatttct ctgtatgatg atgtctctta cttctacagg ttccttgagc ac            412

```

&lt;210&gt; 111

&lt;211&gt; 514

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 111

```

taaattcaca tgcagtctca gagactatth agacaaagtt caagtttagga gcttttagga      60
tgtggggagta aaactttaat gggaggggag ggctggctgc tggaagaagg aagaagccag    120
actggttaga cagtactctt aactcctagc ccagcctacg tgccctgccc ctctggccac    180
tgctgcagac acctgcctta acacacacac ctctaggact ccacagtttt gccttaaagg    240
accttcccaa gtctccctth ccctgtctgg cttctccctt aagaagagag agatacttgt    300
agaattgggt ggggggaatg agcatgaact gtccttccat ttgggatatg ttacattaga    360
gtgagagaga gaataaggag cctttcttat ggaagaaatg ggagaagaga gacagggttc    420
ttttcagcag agtctagtag tttctctgta aggcaaaata atctaaaaag actaacctgc    480
ccaccactc cttatattgc tgtgagattg cccc                                514

```

&lt;210&gt; 112

&lt;211&gt; 489

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 112

```

cggaccatc caagtcatct gattgaagag catgacagaa acaaaatgta ttcaccaagc      60
atthtaggat ttgactthtt cactaaccag ttgacgagca gtgcatttac aaggcactgc    120
caaacaagat gcccttggga gctgtgaggg aaagaggacc tgcgggctta gatcaatctc    180
aattccttht catgccctcc tgcattgctg ctgcgtgggt atthgtctcc ttagccatca    240
ggtacagtht acactacaat gtaagctata ggtggagcat cagcagtgag tgaggccatt    300
cttcatcctt aggatgtggc aatgaaatga tgggtgcaagt tcctthctct tttgtgaatc    360
thtccccca thtctgtht acatgtaacc caacaaatg caatthctag tgctthctgt    420
ccaatcagth cthtctctg agtgagacgt acttggctac agatthctgc cthgtthtgc    480
gacattgtc                                489

```

&lt;210&gt; 113

&lt;211&gt; 416

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

## 40

<400> 113  
gattgggatg gccttagctc ttagccaaac accttcctga caccatgagg gccagcagct 60  
tcttgatcgt ggtgggtgtc ctcatcgctg ggacgctggt tctagaggca gctgtcacgg 120  
gagttcctgt taaagggtcaa gacactgtca aaggccgtgt tccattcaat ggacaagatc 180  
ccgttaaagg acaagtttca gttaaaggtc aagataaagt caaagcgcaa gagccagtca 240  
aaggteccagt ctccactaag cctgggtcct gccccattat cttgatccgg tgcgccatgt 300  
tgaatcccc taaccgctgc ttgaaagata ctgactgccc aggaatcaag aagtgtgtgtg 360  
aaggctcttg cgggatggcc tgtttcgttc cccagtgaag ggagccggtc cttgct 416

<210> 114  
<211> 502  
<212> DNA  
<213> Homo Sapiens

<400> 114  
cccgaaccgt gggcatttgt gaggcccatg gttgagaaat gaataatttc ccaattagga 60  
agtgtgaagca gctgagggtc cttgaggag cttagccaat gtgggagcag cggtttgggg 120  
agcagagaca ctaacgactt cagggcaggg ctctgatatt ccatgaatgt atcaggaaat 180  
atatatgtgt gtgtatgttt gcacacttgt tgtgtgggct gtgagtgtaa gtgtgagtaa 240  
gagctggtgt ctgattgtta agtctaaata tttccttaa ctgtgtggac tgtgatgcca 300  
cacagagtgg tctttctgga gaggttatag gtcactcctg gggcctcttg ggtccccac 360  
gtgacagtgc ctgggaatgt acttattctg cagcatgacc tgtgaccagc actgtctcag 420  
tttcaacttc acatagatgt ccctttcttg gccagttatc ccttcctttt agcctagttc 480  
atccaatcct cactgggtgg gg 502

<210> 115  
<211> 430  
<212> DNA  
<213> Homo Sapiens

<400> 115  
accacaacga cattgccttg ctgaagatcc gttccaagga gggcagggtg gcgcagccat 60  
cccggactat acagaccatc tgccctgccct cgatgtataa cgatccccag tttggcacia 120  
gctgtgagat cactggcttt ggaaaagaga attctaccga ctatctctat ccggagcagc 180  
tgaagatgac tgttgtgaag ctgatttccc accgggagtg tcagcagccc cactactacg 240  
gctctgaagt caccaccaa atgctgtgtg ctgctgacct acagtggaaa acagattcct 300  
gccagggaga ctgaggggga cccctcgtct gttccctcca aggccgcatg actttgactg 360  
gaattgtgag ctggggccgt ggatgtgccc tgaaggacaa gccaggcgtc tacacgagag 420  
tctcacactt 430

<210> 116  
<211> 449  
<212> DNA  
<213> Homo Sapiens

<400> 116  
gggttgccat ccaagtgaag gtcttttctt tgaccaaggg ggacagtcag ttttgcaaaa 60

## 41

ggactctaatacctgttttaa tattgtcttc ctaattggga taatttaatt aacaagattg 120  
 actagaagtgaactgcaac actaacttcc cctgctgtg gtgtgacctg agttggtgac 180  
 acaggccaca gacccagag cttggctttt gaaacacaac tcagggtttt tgtgaaggtt 240  
 ccccgctga gatctttcct cctgggttact gtgaagcctg ttggtttgct gctgtcgttt 300  
 ttgaggaggg cccatggggg taggagcagt tgaacctggg aacaaacctc acttgagctg 360  
 tgcctagaca atgtgaattc ctgtgttgct aacagaagtg gcctgtaagc tcctgtgctc 420  
 cggagggaag catttcctgg taggctttg 449

<210> 117  
 <211> 535  
 <212> DNA  
 <213> Homo Sapiens

<400> 117  
 gctgaaggca gatgtcgtcc caaagacagc tgagaacttc agagccctgt gcaactggtga 60  
 gaagggcttc ggctacaaag gctccacctt ccacaggggtg atcccttcct tcatgtgcca 120  
 ggcgggcgac ttcaccaacc acaatggcac aggcgggaag tccatctacg gaagccgctt 180  
 tcctgacgag aactttacac tgaagcacgt ggggccaggt gtctgttcca tggetaatgc 240  
 tggctctaac accaaccggct ccagttctt catctgcacc ataaagacag actggttgga 300  
 tggcaagcat gttgtgttcg gtcacgtcaa agagggcatg gacgtcgtga agaaaataga 360  
 atctttcggc tctaagagtg ggaggacatc caagaagatt gtcacacag actgtggcca 420  
 gttgagctaa tctgtggcca ggggtgctggc atgggtggcag ctgcaaatgt ccatgcaccc 480  
 aggtggccgc gttgggctgt cagccaaggt gcctgaaacg atacgtgtgc ccaact 535

<210> 118  
 <211> 484  
 <212> DNA  
 <213> Homo Sapiens

<400> 118  
 ggttgaatgt ttgtccttag gataggccta tgtgctagcc cacaagaat attgtctcat 60  
 tagcctgaat gtgccataag actgaccttt taaaatgttt tgagggatct gtggatgctt 120  
 cgttaatttg ttcagccaca atttattgag aaaatattct gtgtcaagca ctgtgggttt 180  
 taatattttt aaatcaaacg ctgattacag ataatagtat ttatataaat aattgaaaaa 240  
 aattttcttt tgggaagagg gagaaaatga aataaatatc attaaagata actcaggaga 300  
 atcttcttta caattttacg tttagaatgt ttaaggttta gaaagaaata gtcaatatgc 360  
 ttgtataaaa cactgttcac tgtttttttt aaaaaaaaaa cttgatttgt tattaacatt 420  
 gatctgctga caaacctgg gaatttgggt tgtgtatgcg aatgtttcag tgcctcagac 480  
 aaat 484

<210> 119  
 <211> 495  
 <212> DNA  
 <213> Homo Sapiens

<400> 119

## 42

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gaacaagcgt cctggggcat ttgctattta cctggagcct tggcatttag acatctttga      60
attccttgat ttaaagaaga acacaggaaa ggaagagcag cgtgccagag atcttttctt      120
tgctcttttg attccggatc tcttcatgaa acgagtggag actaatcagg actgggtctt      180
gatgtgtcca aatgagtgtc ctggtctgga tgaggtttgg ggagaggaat ttgagaaact      240
atatgcaagt tatgagaaac aaggctcgtgt ccgcaaagtt gtaaaagctc agcagctttg      300
gtatgccatc attgagtctc agacggaaac aggcaccccg tatatgctct acaaagattc      360
ctgtaatcga aagagcaacc agcagaacct gggaaccatc aaatgcagca acctgtgcac      420
agaaatagtg gagtacacca gcaaagatga gggtgctggt tgtaatttgg cttccctggc      480
cctgaatatg tatgt                                     495

```

```

<210> 120
<211> 438
<212> DNA
<213> Homo Sapiens

```

```

<400> 120
gcccttgag tcgcgagaaa agggccgtaa ccggaggacc cacgccctg agcctcgcg      60
tgagcggggg ccgcgagcgg caacgcactg gtgaccagac tgtccccacg ccgggaacca      120
agcaggagac gacaggcgag agaggagcca gacagaccct gaaaagaagg acgggttggg      180
gccgggcaca ttgggggtca ccggccgatg gagacaccaa ccgacaggcc ctggctgagg      240
gcagctgcg      300
gggcttattt attaacagga taaccttga atgtagcagc cccgggaggg      360
cggcacaggt cggcgagcgg attcagccgg agggaaggga cggggaagcc gagctccaga      420
gcaacgacca gggccgagga ggtgcctgga gtgcccaccc tgggagacag accccacctc      438
cttgggtagt gagcagtg

```

```

<210> 121
<211> 447
<212> DNA
<213> Homo Sapiens

```

```

<220>
<221> misc_feature
<222> (116)..(116)
<223> n is a, c, g, or t

```

```

<220>
<221> misc_feature
<222> (362)..(362)
<223> n is a, c, g, or t

```

```

<400> 121
ggaactacgg ggcttacagg agcttttgtg tgcttggtag aaactatttc tgttccagtc      60
acattgccat cactcttgta ctgcctgccca ccgcgagga ggctggtgac aggccnaaag      120
gccagtggaa gaaacaccct ttcattctcag agtccactgt ggactggcc acccctcccc      180
agtacagggg tgctgcaggt ggcagagtga atgtcccca tcatgtggcc caactctcct      240
ggcctggcca tctccctccc cagaacaggt gtgcatgggt tattttggag tgtaggtgac      300
ttgtttactc attgaagcag atttctgctt ctttttattt ttataggaat agaggaagaa      360
angtcagatg cgtgcccagc tcttcacccc ccaatctctt ggtggggagg ggtgtaccta      420

```



aatattttatc atatccttgc ccttgag 447

<210> 122  
 <211> 323  
 <212> DNA  
 <213> Homo Sapiens

<400> 122  
 aaattgacca tacaatttca tcctccttca ggggatcaaa aggacggagt ggggggacag 60  
 agactcagat gaggacagag tggtttccaa tgtgttcaat agatttagga gcagaaatgc 120  
 aaggggctgc atgacctacc aggacagaac tttccccaat tacaggggtga ctcacagccg 180  
 cattggtgac tcacttcaat gtgtcatttc cggtgctgt gtgtgagcag tggacacgtg 240  
 aggggggggt ggggtgagaga gacaggcagc tcggattcaa ctaccttaga taatatttct 300  
 gaaaacctac cagccagagg gta 323

<210> 123  
 <211> 499  
 <212> DNA  
 <213> Homo Sapiens

<400> 123  
 gtatcaggct tcaattccat tatgttttaa tgttgtctct gaagatgact tgtgattttt 60  
 ttttcttttt tttaaaccat gaagagccgt ttgacagagc atgctctgcg ttgttggttt 120  
 caccagcttc tgccctcaca tgcacaggga tttaacaaca aaaatataac tacaacttcc 180  
 cttgtagtct cttatataag tagagtcctt ggtactctgc cctcctgtca gtagtggcag 240  
 gatctattgg catattcggg agcttcttag agggatgagg ttctttgaac acagtgaaaa 300  
 tttaaattag taactttttt gcaagcagtt tattgactgt tattgctaag aagaagtaag 360  
 aaagaaaaag cctgttgcca atcttggtta tttctttaag atttctggca gtgtgggatg 420  
 gatgaatgaa gtggaatgtg aactttgggc aagttaaattg ggacagcctt ccatgttcat 480  
 ttgtctacct cttaactga 499

<210> 124  
 <211> 328  
 <212> DNA  
 <213> Homo Sapiens

<400> 124  
 taattttaga ttgccttac aatgtaaatc ttcacattgg agataatatt gggtggacct 60  
 tgcccatctt cactctagcc ttcgtatttg tgaaggactc agccaccttc cttcttcacc 120  
 ccatgcttct caccaaattt ttgttgcac tgagggcact tggataactc aagttgatat 180  
 ttatagctga tcaatctata tgtgtcacag aactatgctg cctaaagtga tcttggtcc 240  
 ttaatggctc ttttggccc ttggatagtt aacagctgag taattetaat ctcttctgtg 300  
 ttttccttgc cttaaccaca aattgtgg 328

<210> 125  
 <211> 489  
 <212> DNA  
 <213> Homo Sapiens

## 44

<400> 125  
gagatacaga acttggtgac ccatgtattg cataagctaa agcaacacag acactcctag 60  
gcaaagtttt tgtttgtgaa tagtacttgc aaaacttgta aattagcaga tgactttttt 120  
ccattgtttt ctccagagag aatgtgctat atttttgtat atacaataat atttgcaact 180  
gtgaaaaaca agttgtgcca tactacatgg cacagacaca aaatattata ctaatatgtt 240  
gtacattcgg aagaatgtga atcaatcagt atgttttttag attgtatttt gccttacaga 300  
aagcctttat tgtaagactc tgatttcctt ttggacttca tgtatattgt acagttacag 360  
taaaattcaa cctttatttt ctaatttttt caacatattg tttagtgtaa agaattttta 420  
tttgaagttt tattatttta taaaaagaa tatttatttt aagaggcatc ttacaaattt 480  
tgccccctt 489

<210> 126  
<211> 503  
<212> DNA  
<213> Homo Sapiens

<400> 126  
gcggcatgtg accatcattg aactggtggg acagccacct caggagggtg ggcgcatccg 60  
ggagcaacag ctgtcagcca acatcatcga ggagctcagg caatttcagc gcctcactcg 120  
ctcctacttc aacatggtgt tgattgacaa gcagggtatt gaccgagacc gctacatgga 180  
acctgtcacc cccgaggaaa tcttcacatt cattgatgac tacctactga gcaatcagga 240  
gttgaccag cgtcgggagc aaagggacat atgcgagtga acttgagcca gggcatggtt 300  
aaagtcaagg gaaaagctcc tctagttagc tgaaactggg acctaataaa aggaggaaat 360  
gttttccac agttctaggg acaggactct gagtggttg agtttgacaa atcctgcagt 420  
gtttccaggc atccttttag gactgtgtaa tagtttcctt agaagctagg tagggactga 480  
ggacaggcct tgggcagtgg gtt 503

<210> 127  
<211> 436  
<212> DNA  
<213> Homo Sapiens

<400> 127  
agactgggag aaaggctgtc cggagggcag accagggtgcc ttgccgcaga gaaaacacca 60  
aagtctcctg ttcgctcata aagaagtttt tgggatggga gagaatccag accatcttgg 120  
ggcagccagg cccttgcctt catTTTTtaca gaggtagcac aactgattcc aacacaaaac 180  
cccttccctt ttttaaaatg atttctgttc taatgccata gatcaaaggc ctcaaaaacc 240  
attgtgtgtt tcctctttga agcaatgaca agcactttac tttcacgggtg gtttttgttt 300  
tttcttattg ctgtggaacc tcttttgag gaggttaaag gcgtgtttta cttgtttttt 360  
taagagtgtg tgatgtgtgt tttgtagatt tcttgacagt gctgtaatac agacggcaat 420  
gcaatagcct atttaa 436

<210> 128  
<211> 497  
<212> DNA  
<213> Homo Sapiens

<400> 128  
 cctgccctct agttgggtctt gggctttgat ctcttccaac ctgccagtc acagaaggag 60  
 gaatgactca aatgccccaa accaagaaca cattgcagaa gtaagacaaa catgtatatt 120  
 tttaaatggt ctaacataag acctgttctc tctagccatt gatttaccag gctttctgaa 180  
 agatctagt gttcacacag agagagagag agtactgaaa aagcaactcc tcttcttagt 240  
 cttaataatt tactaaaatg gtcaactttt cattatcttt attataataa acctgatgct 300  
 tttttttaga actccttact ctgatgtctg tatatgttgc actgaaaagg ttaatatatta 360  
 atgttttaat ttattttgtg tggtaagtta attttgattt ctgtaatgtg ttaatgtgat 420  
 tagcagttat tttccttaat atctgaatta tacttaaaga gtagtgagca atataagacg 480  
 caatgtgtt tttcagt 497

<210> 129  
 <211> 321  
 <212> DNA  
 <213> Homo Sapiens

<400> 129  
 gtttgatgg tggaaggctt cattttattg agatttttaa gatacatgca aagggttgga 60  
 aatagaacct ctaggcacc cctcagtggt ggggtggctg agagttaaag acagtgtggc 120  
 tgcagtagca tagaggcgcc tagaaattcc acttgcaccg tagggcatgc tgataccatc 180  
 ccaatagctg ttgccattg acctctagt gtaggtttct agaatactgg tccattcatg 240  
 agatattcaa gattcaagag tattctcact tctgggttat cagcataaac tggaatgtag 300  
 tgtcagagga tactgtggct t 321

<210> 130  
 <211> 553  
 <212> DNA  
 <213> Homo Sapiens

<400> 130  
 tttgcctgca gtttcttgtg tagatttgaa aattgtatac caatgtgttt tctgtagact 60  
 ctaagataga ctgcactttg tttagaaaaa aaactgaaga tgaaatatat attgtaaaga 120  
 agggatatta agaactcttag ataacttctt gaaaaagatg gcttatgtca tcagtaaagt 180  
 acctttatgt tatgaggata taatgtgtgc tttattgaat tagaaaatta gtgaccatta 240  
 ttcacagggt gacaaatggt gtctctgttaa tttataggag ttttttgggg atgtggaggt 300  
 agttgggtag aaaaattatt agaacattca cttttgttaa cagtatttct cttttattct 360  
 gttatatagt ggatgatata cacagtggca aaacaaaagt acattgctta aaatatatag 420  
 tgaaaaatgt cactatatct tcccatTTaa cattgttttt gtatatggg ttagatttc 480  
 tgacatcaaa acttggaccc ttggaaaaca aaagtTTTaa ttaaaaaaa tccttgtgac 540  
 ttacaatttg cac 553

<210> 131  
 <211> 419  
 <212> DNA  
 <213> Homo Sapiens

## 46

<400> 131  
gagtcggaga tgatgcagca cacacacaat tccccagccc agtgatgctt gtgttgacca 60  
gatgttcctg agtctggagc aagcaccag gccagaataa cagagcttcc ttagttgggtg 120  
aagacttaaa catctgcctg aggtcaggag gcaatttgcc tgccttgtag aaaagctcag 180  
gtgaaagact gagatgaatg tctttcctct ccctgcctcc caccagactt cctcctggaa 240  
aacgctttgg tagatttggc caggagcttt cttttatgta aattggataa atacacacac 300  
catacactat ccacagatat agccaagtag atttgggtag aggatactat ttccagaata 360  
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<210> 132  
<211> 414  
<212> DNA  
<213> Homo Sapiens

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atctgacccc tgtccgtctc cttgtccctg cttcatgttt gggggccttt ctttaactgc 180  
cttctgggt tagctcagat ggcagatgag agtgtagtca agggcctggg cacaggaggg 240  
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agcttgggtc cctttcccta gctccctggt gggatgaatgc cacctcctga gacccacc 360  
tcttgaatt aaaattgttg gtcactgggg aaagcctgag tttgcaacca gttg 414

<210> 133  
<211> 419  
<212> DNA  
<213> Homo Sapiens

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tgccttgctt tgtgacttcc aagaacgagt gtctctggac cgacatgctc tccaatttcg 120  
gttaccctgg ctaccagtcc aaacactacg cctgcatccg gcagaagggc ggctactgca 180  
gctggtaccg aggatggggc cccccgata aaagcatcat caatgccaca gaccctgag 240  
cgccagaccc tgcccacct cacttccctc ccttcccgct gagcttccct tggacactaa 300  
ctcttcccag atgatgacaa tgaaattagt gcctgttttc ttgcaaattt agcacttgga 360  
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<212> DNA  
<213> Homo Sapiens

<400> 134  
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cccaagtgtt tggcttctgg ctacctaagg ttaacatgtc actagagtat ttttatgaga 180  
gacaaacatt ataaaaatct gatggcaaaa gcaaaacaaa atggaaagta ggggaggtgg 240

## 47

atgtgacaac aacttccaaa ttggctcttt ggaggcgaga ggaaggggag aacttggaga 300  
 atagtttttg ctttgggggt agaggcttct tagattctcc cagcatccgc ctttcccttt 360  
 agccagtctg ctgtcctgaa acccagaagt gatggagaga aaccaacaag agatctcgaa 420  
 ccctgtctag aaggaatgta tttgttgcta aatttcgtag cactgtttac agttttcttc 480  
 catgttattt atg 493

<210> 135  
 <211> 567  
 <212> DNA  
 <213> Homo Sapiens

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 gactgccctt agaaattcta gcctggtttg gagatactaa ctgctctcag agaaagtagc 180  
 tttgtgacat gtcattgaacc catgtttgca atcaaagatg ataaaataga ttcttatttt 240  
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 tgtattaaat gtgaatttta agaaataata tttatatattc tgtaaatgta aactgtgaag 480  
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<210> 136  
 <211> 479  
 <212> DNA  
 <213> Homo Sapiens

<220>  
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 <223> n is a, c, g, or t

<400> 136  
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 aaggctttgt tcgaaagaat ggtgtcaatg aagccaaaat agatgagatc aagaatgaca 180  
 atgtccaaga cacagcagaa cagaaagttc aactgcttcg taattggcat caacttcatg 240  
 gaaagaaaaga agcgtatgac acattgatta aagatctcaa aaaagccaat ctttgtactc 300  
 ttgcagagaa aattcagact atcatcctca aggacattac tagtgactca gaaaattcaa 360  
 acttcagaaa tgaaatccaa agcttgggtc agagtgaata acaacaaatt cagttctgag 420  
 tatatgcaat tagtgtttga naagattctt aatagctggc tgtaataact gcttggttt 479

<210> 137  
 <211> 490  
 <212> DNA  
 <213> Homo Sapiens

## 48

<400> 137  
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 tcattcctga gcaacttgca cttgaggaat ggtgaactgg tcatccatga aaaagggttt 180  
 tactacatct attcccaaac atactttcga tttcaggagg aaataaaaga aaacacaaag 240  
 aacgacaaac aaatggtcca atatatttac aaatacacia gttatcctga ccctatatgt 300  
 ttgatgaaaa gtgctagaaa tagttgttgg tctaaagatg cagaatatgg actctattcc 360  
 atctatcaag ggggaatatt tgagcttaag gaaaatgaca gaatttttgt ttctgtaaca 420  
 aatgagcact tgatagacat ggaccatgaa gccagttttt tcggggcctt tttagttggc 480  
 taactgacct 490

<210> 138  
 <211> 248  
 <212> DNA  
 <213> Homo Sapiens

<400> 138  
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 cgcaacaatc catctctcaa gtagtgtatc acagtagtag cctccagggt tccttaaggg 180  
 acaacatcct taagtcaaaa gagagaagag gcaccactaa aagatcgag tttgctgggt 240  
 gcagtggc 248

<210> 139  
 <211> 405  
 <212> DNA  
 <213> Homo Sapiens

<220>  
 <221> misc\_feature  
 <222> (64)..(64)  
 <223> n is a, c, g, or t

<220>  
 <221> misc\_feature  
 <222> (68)..(68)  
 <223> n is a, c, g, or t

<400> 139  
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 ccgntanct gacggtggcc accgtgttcc ggggccgcat gtccatgaag gaggtggacg 120  
 agcagatgct ggccatccag agcaagaaca gcagctactt cgtggagtgg atccccaaca 180  
 acgtgaaggt ggccgtgtgt gacatccgc cccgcggcct caagatgtcc tccaccttca 240  
 tcgggaacag cacggccatc caggagctgt tcaagcgcgt ctccgagcag ttcacggcca 300  
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<210> 140  
 <211> 407  
 <212> DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 140

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aaagcagtgt gcttcaaagg catcagacga tgaaagcaac ataccacaac taggagttat	120
ttctcaaact taaatgtcct ctgggaatcc agacttaaaa ataagagcaa acttaacaca	180
ctatccattt tcgagcaaac ttaaccact atatccattt tgctcatgtg ttttatgcaa	240
ccagctttcc atcaaactct caatccttga atccaggtaa aagggttaatt atcctaggat	300
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gtttgtatcc tgtttttagtt tataaagcac tttcacatac attatgg	407

&lt;210&gt; 141

&lt;211&gt; 518

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 141

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ttatttgaga aggaacaact tggagttcat ctataacaag actggttggg ccatggtgtc	120
tctgtgtata gtctttgcta tgacttctgg ccagatgtgg aaccatatcc gtggacctcc	180
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ggttcttcta aatgaagcag caacttcgaa aggcgatgtt ggaaaaagac ggataatttg	360
cctagtggga ttgggcctgg tggcttctct cttcagtttt ctactttcaa tatttcgttc	420
caagtaccac ggctatcctt atagtgatct ggactttgag tgagaagatg tgatttggac	480
catggcactt aaaaactcta taacctcagc cttttaat	518

&lt;210&gt; 142

&lt;211&gt; 443

&lt;212&gt; DNA

&lt;213&gt; Homo Sapiens

&lt;400&gt; 142

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gaaccacac aatggacaag tgagctacat tcatgggagc agccaggctc agtttctggc	120
agaatcacac attattctgg tactgaatgc cgctatcacc atggggatgg ttcttctaaa	180
tgaagcagca acttcgaaag gcgatgttgg aaaaagacgg ataatttgcc tagtgggatt	240
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ctatccttat agctttttta ttaaatgaag ccaagtggga tttgcataaa gtgaatgttt	360
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